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cagcctgacatctgaagactctgcggtctatttctgtgcaagagtggtgtactatagtaactcttactggtacttcgatgtctggggcac agggaccaeggtcaccgtctcttctgatcaatccaactctgaagaagcaaagaagaggaggccaaaaaaggaggaagccaaga aatctaacagcglcgacattgttetgactcagtetecagecaccetgtetgtgactccaggagatagagtetetettteetgeagggee agccagagtattagcgactacttacactggtatcaacaaaaatcacatgagtctccaaggcttctcatcaaatatgcttcccattccatc5 tctgggatcccctccaggttcagtggcagtggatcagggtcagatttcactctcagtatcaacagtgtggaacctgaagatgttggaa ittattactgtcaacatggtcacagctttccgtggacgttcggtggaggcaccaagctggaaatcaaacggggtggcggtggctcg ggcggaggtgggtcgggtggcggatetcagatecagttggtgcaatetggacetgagetgaagaagcetggagagacagt caggatctcctgcaaggcttctgggtatgccttcacaactactggaatgcagtgggtgcaagagatgccaggaaagggtttgaagt 10 ggattggctggataaacacccactctggagtgccaaaatatgtagaagacttcaaggacggtttgccttcttttggaaacctctgc caa cact g cat at tta caga taa g caa cct caa ag at gag ga cac g g ctac g tat tt ct g t g t g ag at ccg g gaa t g g taa ct at g a cac g cat at tt ct g t g t g ag at ccg g g a at g g t a ct at g a cac g cat at tt ct g t g t g ag at ccg g g a at g g t a ct at g a cac g cat at tt ct g t g t g ag at ccg g g a at g g t a ct at g a cac g cat at tt ct g t g t g ag at ccg g g a at g g t a ct at g a cac g cat at tt ct g t g t g ag at ccg g g a at g g t a ct at g a cac g cat at tt ct g t g t g ag at ccg g g a at g g t a ct at g a cac g ccctggcctactttgcttactggggccaagggacaetggtcactgtctctgatcaggagcccaaatcttctgacaaaactcacacatceccaccgtccccagcacctgaactcctggggggatcgtcagtcttcctcttccccccaaaacccaaggacaccctcatgatctcccg gacccct gagg tcacat gcgt ggt ggt ggacgt gagccac gaag accct gagg tcaa gttcaact ggt ac gt ggacg gcgt ggacgt gaggacgt gag15 ggtgcataatgccaagacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcacca ggaclggctgaatggcaaggagtacaagtgcaaggtctccaacaaagccctcccaguccccatcgagaaaacaatctccaaagc caa aggg cag coccgaga acca caggt gia caccet g coccat cocgg gat gag ctgac caa gaa ccagg to agcet gacctgcctggtcaaaggcttctatcccagcgacatcgccgtggagtgggagagcaatgggcagccggagaacaactacaagaccacg cctcccgtgctggactccgacggctccttcttcctctacagcaagctcaccgtggacaagagcaggtggcagcaggaacgtct 20 tctcatgctccgtgatgcatgaggctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaatgatctaga

# 2H7-antiCD40 scFv MTH (SSS) MTCH2WTCH3 (2H7-40.2.220Ig) (amino acid sequence) (SEQ ID NO: )

MDFOVOIFSFLLISASVIIARGQIVLSOSPAILSASPGEKVTMTCRASSSVSYMHWY 2.5 QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCOOWS FNPPTFGAGTKLELKGGGGGGGGGGGGGGGGSOAYLOOSGAELVRPGASVKMSCK ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDOSNSEEAK KEEAKKEEAKKSNSVDIVLTOSPATLSVTPGDRVSLSCRASOSISDYLHWYOOKSH ESPRLLIKYASHSISGIPSRFSGSGSGSDFTLSINSVEPEDVGIYYCOHGHSFPWTFGG

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EDTATYFCVRSGNGNYDLAYFAYWGQGTLVTVSDQEPKSSDKTHTSPPSPAPELL GGSSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEVHNAKTK PREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAKGQPRE PQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPPVLDSD GSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTOKSLSLSPGK

GSFFLYSKLIVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSPGK

5B9 VH (minus the leader) (nucleotide sequence) (SEQ ID NO:\_\_\_)

15 caggtgeagetgaageagicaggacetgggeetragtgeagtoctcacagageetgtecateacetgeacagtetetggttteteatta. actacetatgetgtacaetgggttegecagictecaggaaagggtetggagtggetgggagtgatatggagtggtggaateacaga ctataatgeagettteatatecagactgageateaceaaggaegattecaagagecaagtttietttaaatgaacagtetgeaaceta atgacacagecatttattactgtgccagaaatgggggtgataactaccettattactatgetatggaetactggggtcaaggaacetea gteacegteteetea

20

5B9 VH (includes leader peptide) (amino acid sequence) (SEQ ID NO: \_\_\_\_)

MAVLGLLFCLVTFPSCVLSQVQLKQSGPGLVQSSQSLSITCTVSGFSLTTYAVHWV
RQSPGKGLEWLGVIWSGGITDYNAAFISRLSITKDDSKSQVFFKMNSLQPNDTAIY
YCARNGGDNYPYYYAMDYWGOGTSVTVSS

25

5B9 VH (no leader peptide) (amino acid sequence) (SEQ ID NO:\_\_)
QVQLKQSGPGLVQSSQSLSITCTVSGFSLTTYAVHWVRQSPGKGLEWLGVIWSGGI
TDYNAAFISRLSITKDDSKSQVFFKMNSLQPNDTAIYYCARNGGDNYPYYYAMDY
WGQGTSVTVSS

## 5B9 VL (nucleotide sequence) (SEQ ID NO:\_\_\_)

## 5B9 VL (amino acid sequence) (SEQ ID NO: )

MRFSAQLLGLLVLWIPGSTADIVMTQAAFSNPVTLGTSASISCRSSKSLLHSNGITY

LYWYLQKPGQSPQLLIYQMSNLASGVPDRFSSSGSGTDFTLRISRVEAEDVGVYYC

AQNLELPLTFGAGTKLELKR

### 5B9 scFv (nucleotide sequence) (SEQ ID NO: )

# 25 5B9 scFv (amino acid sequence) (SEQ ID NO:\_\_\_)

MRFSAQLLGLLVLWIPGSTADIVMTQAAFSNPVTLGTSASISCRSSKSLLHSNGITY LYWYLQKPGQSPQLLIYQMSNLASGVPDRFSSSGSGTDFTLRISRVEAEDVGVYYC AQNLELPLTFGAGTKLELKRGGGGSGGGGSGGGGSSQVQLKQSGPGLVQSSQSLS ITCTVSGFSLTTYAVHWVRQSPGKGLEWLGVIWSGGITDYNAAFISRLSITKDDSK

30 SQVFFKMNSLQPNDTAIYYCARNGGDNYPYYYAMDYWGQGTSVTVSS

### 5B9 scFv-hmtIgG1-hCD80 (nucleotide sequence) (SEQ ID NO:

ggctgcattctccaatccagtcactcttggaacatcagcttccatctcctgcaggtctagtaagagtctcctacatagtaatggcatca 5 ctt att t g t att g g t at et g cag a age cag g c cag et c c t cag et c c t g at t t at eag at g t c cag ag et c cag ag et cag at g t cag at g t cag ag et cag ag etggttcagtagcagtgggtcaggaactgatttcacactgagaatcagcagagtggaggctgaggatgtgggtgtttattactgtgctc aaaatctagaacttccgctcacgttcggtgctgggaccaagctggagctgaaacggggtggcggtggctcgggcggtggtggt cgggtggcggcggategtcacaggtgcagctgaagcagtcaggacctggcctagtgcagtcctcacagagcctgtccatcacct gcacagtctctggtttctcattaactacctatgctgtacactgggttcgccagtctccaggaaagggtctggagtggctgggagtgat 10 atggagtggtggaatcacagactataatgcagctttcatatccagactgagcatcaccaaggacgattccaagagccaagtttcttt aaaatgaacagtetgcaacetaatgacacagccatttattactgtgccagaaatgggggtgataactacccttattactatgctatgga ctactggggtcaaggaacctcagtcaccgtctcctctgatctggagcccaaatcttctgacaaaactcacacaagcccaccgagcc cagcacctgaactcctgggggggatcgtcagtcttcctcttcccccaaaacccaaggacaccctcatgatctcccggacccctgag gtcacatgcgtggtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataat 15 gccaagacaaagcegcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggct gaatggcaaggagtacaagtgcaaggtetecaacaaageceteccagececategagaaaaecatetecaaagecaaaggge agccccgagaaccacaggigtacaccctgccccatcccgggatgagctgaccaagaaccaggtcagcctgacctgctcgtca aaggettetateeeagegacategeegtggagtgggagageaatgggeageeggagaacaactacaagaceaegeeteeegtg ctggactccgacggctccttcttcctctacagcaagctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctc 20 cgtgatgcatgaggctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaagcggatccttcgaacctgctcc aggaggaatgagagttgagaagggaaagtgtacgccctgtataaatcgatactcgag

## 5B9 scFv-hmtIgG1-hCD80 (amino acid sequence) (SEQ ID NO:

25 MRFSAQLLGLVLWIPGSTADIVMTQAAFSNPVTLGTSASISCRSSKSLLHSNGITY
LYWYLQKPGQSPQLLIYQMSNLASGVPDRFSSSGSGTDFTLRISRVEAEDVGVYYC
AQNLELPLTFGAGTKLELKRGGGSGGGGGGGGGSSQVQLKQSGPGLVQSSQSLS
ITCTVSGFSLTTYAVHWVRQSPGKGLEWLGVIWSGGITDYNAAFISRLSITKDDSK
SQVFFKMNSLQPNDTAIYYCARNGGDNYPYYYAMDYWGQGTSVTVSSDLEPKSS
30 DKTHTSPPSPAPELLGGSSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNW
YVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPA

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ENNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTOKSLS LSPGKADPSNLLPSWAITLISVNGIFVICCLTYCFAPRCRERRNERLRRESVRPV

# 2e12 scFv WTH CH2 CH3 (2e12 scFv-WthIgG-CD80) (nucleotide sequence) (SEO ID NO: )

aagcttatggattttcaagtgcagattttcagcttcctgctaatcagtgcttcagtcataatgtccagaggagtcgacattgtgctcaccc aatctccagettetttggetgtgtetetaggtcagagggccaccateteetgcagaggcagtgaaagtgttgaatattatgtcacaagtt taatgcagtggtaccaacagaaaccaggacagccacccaaactcctcatctctgctgcatccaacgtagaatctggggtccctgcc 10 tegggtggeggeggateteaggtgeagetgaaggagteaggaeetggeetggtggegeeeteaeagageetgteeateaeatge accgtctcagggttctcattaaccggctatggtgtaaactgggttcgccagcctccaggaaagggtctggagtggctgggaatgat atggggtgatggaagcacagactataattcagctctcaaatccagactgagcatcaccaaggacaactccaagagccaagttttett aaaaatgaacagtctgcaaactgatgacacagccagatactactgtgccagagatggttatagtaactttcattactatgttatggactgeaectgaactcetggggggaecgtcagtettcetettcecccaaaacccaaggacaccetcatgatctcceggacccetgaggt cacatgcgtggtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctga atggcaaggagtacaagtgcaaggtctccaacaaagccctcccagccccatcgagaaaaccatctccaaagccaaagggcag ggettetateceagegacategeegtggagtgggagageaatgggeageeggagaacaactacaagaceaegeeteeegtget ggacteegacggcteettelteetetaeageaageteaeegtggacaagageaggtggeageaggggaaegtetteteatgeteegtgatgcatgaggctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaagcggatccttcgaacctgctccc atectgggccattacettaateteagtaaatggaatttttgtgatatgetgectgacetaetgetttgececaagatgeagagagaa ggaggaatgagagattgagaagggaaagtgtacgccctgtataaatcgat

# 2e12 scFv WTH CH2 CH3 (2e12 scFv-WthIgG-CD80) (amino acid sequence) (SEQ ID NO: )

MDFQVQIFSFLLISASVIMSRGVDIVLTQSPASLAVSLGQRATISCRASESVEYYVTS 30 LMOWYOOKPGOPPKLLISAASNVESGVPARFSGSGSGTDFSLNIHPVEEDDIAMYF CQQSRKVPWTFGGGTKLEIKRGGGGSGGGGGGGGGGQVQLKESGPGLVAPSOSLS ITCTVSGFSLTGYGVNWVROPPGKGLEWLGMIWGDGSTDYNSALKSRI SITKDNS

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KSQVFLKMNSLQTDDTARYYCARDGYSNFHYYVMDYWGQGTSVTVSSDLEPKS CDKTHTCPPCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFN WYVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALP APIEKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQ PENNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSL

SLSPGKADPSNLLPSWAITLISVNGIFVICCLTYCFAPRCRERRNERLRRESVRPV

2H7-human IgE Fc (CH2-CH3-CH4) (nucleotide sequence) (SEO ID NO:

aagettgeegeeatggatttteaagtgeagatttteagetteetgetaateagtgetteagteataattgeeagaggaeaaattgttetet cccagtctccagcaatcctgtctgcatctccaggggagaaggtcacaatgacttgcagggccagctcaagtgtaagttacatgcact ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccacccacgttcggtgctgggaccaagctggagctgaaaggtggcggtggctcgggcggtggttggatctggaggaggtg ggageteteaggettatetacageagtetggggetgagetggtgaggeetggggeeteagtgaagatgteetgeaaggettetgge tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat ggtgatactteetacaateagaagtteaagggeaaggeeacaetgactgtagacaaateeteeageacageetacatgeageteag cageetgacatetgaagaetetgeggtetatttetgtgeaagagtggtgtaetatagtaaetettaetggtaettegatgtetggggeae agggaceaeggteaeegtetetgateaegtetgeteeagggactteaeeeegeeeaeegtgaagatettaeagtegteetgegaeg ggacgggcaggtcatggacgtggacttgtccaccgcctctaccacgcaggagggtgagctggcctccacacaaagcgagctca ccctcagccagaagcactggctgtcagaccgcacctacacctgccaggtcacctatcaaggtcacacctttgaggacagcaccaa gaaccactccaccagaaaggaggagaagcagcgcaatggcacgttaaccgtcacgtcaccctgccggtggcacccgagact  $\tt cggcccgcgtgctgccccggaagtctatgcgtttgcgacgccggagtggccgggagccgggacaagcgcaccctcgcctgc$ etgatecagaactteatgeetgaggacateteggtgeagtggetgeacaaegaggtgeageteeeggaegeeeggeaeageaeg acgcagccccgcaagaccaagggctccggcttcttcgtcttcagccgcctggaggtgaccagggccgaatgggagcagaaaga 

30 aatctaga

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2H7 scFv IgE (CH2-CH3-CH4) (amino acid sequence) (SEQ ID NO:\_\_)

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS
FNPPTFGAGTKLELKGGGGSGGGGGGGGSGGGSQAYLQQSGAELVRPGASVKMSCK
ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSDHVCSRDFTP
PTVKILQSSCDGGGHFPPTIQLLCLVSGYTPGTINTTWLEDGQVMDVDLSTASTTQE
GELASTQSELTLSQKHWLSDRTYTCQVTYQGHTFEDSTKKCADSNPRGVSAYLSR
PSPFDLFIRKSPTITCLVVDLAPSKGTVNLTWSRASGKPVNHSTRKEEKQRNGTLTV
TSTLPVGTRDWIEGETYQCRVTHPHLPRALMRSTTKTSGPRAAPEVYAFATPEWP
GSRDKRTLACLIQNFMPEDISVQWLHNEVQLPDARHSTTQPRKTKGSGFFVFSRLE

## 2H7 scFv MH (SSS) MCH2WTCH3 (nucleotide sequence) (SEQ ID NO:\_\_\_)

VTRAEWEQKDEFICRAVHEAASPSQTVQRAVSVNPGK

15 aagettgccgccatggattttcaagtgcagattttcagcttcctgctaatcagtgcttcagtcataattgccagaggacaaattgttctctcccagtetecageaatectgtetgeatetecagggggagaaggteacaatgaettgeagggeeageteaagtgtaagttacatgeact ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcggggggtggtggtctggaggaggtg 20 ggagctctcaggcttatctacagcagtctggggctgagctggtgaggcctggggcctcagtgaagatgtcctgcaaggcttctggc tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat cagcet gacatet gaag actet geggtet at titet gtge aag ag tigtget act at ag taactet ta et gegtaet te gat get each gat gegen actet gat get gegen actet gat get gegen actet gat get gegen actet gat gegen gegenagggaccacggtcaccgtctcttctgatcaggagcccaaatcttctgacaaaactcacacatccccacgtccccagcacctgaac 25 tectggggggategteagtetteetetteecccaaaaacccaaggacacceteatgateteeeggacccetgaggteacatgegtg gtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaag ccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaagga gtacaagtgcaaggtctccaacaaagccctcccagccccatcgagaaaacaatctccaaagccaaagggcagcccgagaac cacaggtgtacaccetgccccatcccgggatgagetgaccaagaaccaggtcagcetgacetgcctgctggtcaaaggettetatcc 30 cagegacategeegtggagtgggagageaatgggeageeggagaacaactacaagaccaegeeteeegtgetggacteegae ggctccttcttcctctacagcaagctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatga ggctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaatgatctaga

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2H7 scfv MH (SSS) MCH2WTCH3 (amino acid sequence) (SEQ ID NO:\_\_\_)
MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
QQKPGSSPKPWIYAPSNI.ASGVPARFSGSGSGTSYSLITISRVEAEDAATYYCQQWS
FNPPTFGAGTKLELKDGGGSGGGGGGGGGSQGAYLQQSGAELVRPGASVKMSCK
ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSSDK
THTSPPSPAPELLGGSSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV
DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIE
KTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPEN
NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS
PGK

### 5B9 scFv MTHWTCH2CH3 (nucleotide sequence) (SEQ ID NO: )

15 a agett g cegecat g aggtte tet gete agette t g g g g et get g t g et e t g g at e cet g g at eggetgeattetecaatecagteactettggaacateagettecateteetgeaggtetagtaagagteteetaeatagtaatggeatea cttatttgtattggtatctgcagaagccaggccagtctcctcagctcctgatttatcagatgtccaaccttgcctcaggagtcccagaca ggttcagtagcagtgggtcaggaactgatttcacactgagaatcagcagagtggaggctgaggatgtgggtgtttattactgtgctc aaaatetagaacticegeteacgtteggtgetgggaccaagetggagetgaaacggggtggeggtggeteggeggtggtggtggt 20 cgggtggcggcggatcgtcacaggtgcagctgaagcagtcaggacctggcctagtgcagtcctcacagagcctgtccatcacct gcacagtctctggtttctcattaactacctatgctgtacactgggttcgccagtctccaggaaagggtctggagtggctgggagtgat atggagtggtggaatcacagactataatgcagctttcatatccagactgagcatcaccaaggacgattccaagagccaagttttcttt aaaatgaacagtctgcaacctaatgacacagccatttattactgtgccagaaatgggggtgataactacccttattactatgctatgga ctactggggtcaaggaacctcagtcaccgtctcctctgatcaggagcccaaatcttctgacaaaactcacacatccccaccgtcccc 25 agcacetgaacteetggggggacegteagtetteetetteececeaaaacceaaggacacecteatgateteeeggacecetgag gtcacatgcgtggtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataat gccaagacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggct gaalggcaaggagtacaagtgcaaggtetecaacaaageceteccagececategagaaaacaatetecaaagecaaaggge 30 aaggettetateecagegacategeegtggagtgggagagcaatgggcageeggagaacaactacaagaccaegceteeegtg ctggactccgacggctccttcttcctctacagcaagctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctc egtgatgcatgaggetetgeacaaccactacaegcagaagageeteteeetgteteegggtaaatgatetaga

5B9 scFv MTHWTCH2CH3 (amino acid sequence) (SEQ ID NO:\_\_)

MRFSAQLLGLLVLWIPGSTADIVMTQAAFSNPVTLGTSASISCRSSKSLLHSNGITY
LYWYLQKPGQSPQLLIYQMSNLASGVPDRFSSSGSGTDFTLRISRVEAEDVGVYYC

5 AQNLELPLTFGAGTKLELKRGGGGSGGGGSGGGGSSQVQLKQSGPGLVQSSQSLS
ITCTVSGFSLTTYAVHWVRQSPGKGLEWLGVIWSGGITDYNAAFISRLSITKDDSK
SQVFFKMNSLQPNDTAIYYCARNGGDNYPYYYAMDYWGQGTSVTVSSDQEPKSS
DKTHTSPPSPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNW
YVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPA
10 PIEKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQP
ENNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLS
LSPGK

#### Human IgG1 hinge mutations

2H7 scFv- MTH (CSS) WTCH2CH3 (nucleotide sequence) (SEQ ID NO: )

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aagettgeegeeatggattiteaagtgeagattiteagetieetgetaateagtgetteagteataattgeeagaggaeaaattgttetet cccagtctccagcaatcctgtctgcatctccaggggagaaggicacaatgacttgcagggccagctcaagtgtaagttacatgcact ggtaccagcagaagccaggatcctcccccaaacctggatttatgccccatccaacctggettctggagtcctgctcgcttcagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgtteggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggageteteaggettatetaeageagtetggggetgagetggtgaggeetgggggeeteagtgaagatgteetgeaaggettetgge tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat cagoctgacatctgaagactctgoggtctatttctgtgcaagagtggtgtactatagtaactcttactggtacttcgatgtctggggcac agggaccacggtcaccgtctcttctgatcaggagcccaaatcttgtgacaaaactcacacatccccacgtccccaggacctgaac tectggggggaccgtcagtettectettecccccaaaacccaaggacacctcatgateteccggacccctgaggtcacatgcgtg gtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacgtggaggtgcataatgccaagacaaag ccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaagga gtacaagtgcaaggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaagccaaagggcagcccgagaac cagcgacatcgccgtggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgac

ggeteettetteetetacageaageteacegtggacaagageaggtggcageaggggaacgtetteteatgeteegtgatgeatga ggetetgcacaaccactacacgeagaagageeteteeetgteteegggtaaatgatetaga

# 2H7 scFv- MTH (CSS) WTCH2CH3 (amino acid sequence) (SEQ ID NO:\_\_\_)

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS FNPPTFGAGTKLELKDGGGSGGGGSGGGGSGQAYLQQSGAELVRPGASVKMSCK ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSCDK THTSPPSPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIE KTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWSNGQPEN NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS PGK

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# 2H7 scFv- MTH (SCS) WTCH2CH3 (nucleotide sequence) (SEQ ID NO:\_\_\_)

ggetcettetteetetacageaageteacegtggacaagageaggtggcageaggggaacgtettetcatgetcegtgatgcatga ggetetgeacaaccactacaegcagaagageeteteeetgteteegggtaaatgatetaga

2H7 scFv- MTH (SCS) WTCH2CH3 (amino acid sequence) (SEQ ID NO: ) 5 MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCOOWS FNPPTFGAGTKLELKDGGGSGGGGGGGGGGSOAYLOOSGAELVRPGASVKMSCK ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS 10 TAYMOLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDOEPKSSDK THTCPPSPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWY VDGVEVHNAKTKPREEOYNSTYRVVSVLTVLHODWLNGKEYKCKVSNK ALP API EKTISKAKGOPREPOVYTLPPSRDELTKNOVSLTCLVKGFYPSDIAVEWESNGOPE NNYKTTPPVLDSDGSFFLYSKLTVDKSRWOOGNVFSCSVMHEALHNHYTOKSLSL 15 SPGK

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2H7 scFv- MTH (SSC) WTCH2CH3 (nucleotide sequence) (SEQ ID NO: \_\_\_\_)

aagettgeegecatggatttteaagtgeagatttteagetteetgetaateagtgetteagteataattgeeagaggaeaaattgtetet cccag to tecage a at cct giot geate to cag gg gag a aggic a cant gact t geag gg ccage to a agt gt a agt tacat geact.ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggageteteaggettatetacageagtetggggetgagetggtgaggeetggggeeteagtgaagatgteetgeaaggettetgge tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat 25 ggtgataetteetaeaateagaagtteaagggeaaggeeacaetgaetgtagacaaateeteeageacageetaeatgeageteag cagcctgacatetgaagactetgeggtetatttetgtgeaagagtggtgtactatagtaactettactggtacttegatgtetggggeae agggaccacggtcaccgtctcttctgatcaggagcccaaatcttctgacaaaactcacacatccccaccgtgcccagcacctgaac tcctggggggaccgtcagtcttcctcttccccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcacatgcgtg gtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaag ccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaagga gtacaagtgcaaggtetecaacaaageceteccagececcategagaaaacaatetecaaagecaaagggcagecegagaac

cacaggtgtacacctgeccccatcccgggatgagctgaccaagaaccaggteagctgacctgcctggtcaaaggettetatcc cagcgacatcgccgtggagtgggagagcaatgggcagcgggagaacaactacaagaccacgcctcccgiggtggactccgac ggctccttcttctctctacagcaagctcaccgtggacaagagcaggtggcagcagggggaacgtcttctcatgctccgtgatgcatga ggctctgcacaaccactacacgcagaagagcctctccctgtctccggggaaatgatctaga

2H7 scFv- MTH (SSC) WTCH2CH3 (amino acid sequence) (SEQ ID NO:\_\_)

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
QQKPGSSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS
FNPPTFGAGTKLELKDGGGSGGGGSGGGGSSQAYLQQSGAELVRPGASVKMSCK

10 ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSSDK
THTSPPCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVKFNWY
VDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPI
EKTISKAKGQPREPQVYTLPPSRDELITNQVSLTCLVKGFYPSDLAVEWESNGQPE

15 NNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSL

HIgGMHcvs1 (nucleotide sequence) (SEQ ID NO:\_\_) gtt gtt gat cag gag ccc aaa tct tct gac aaa act cac aca tg

SPGK

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HIgGMHcys2 (nucleotide sequence) (SEQ ID NO:\_\_\_)
gtt gtt gat cag gag ccc aaa tct tgt gac aaa act cac aca tct cca ccg tgc

HIGGMHcys3 (nucleotide sequence) (SEQ ID NO:\_\_\_)

25 gtt gtt gat cag gag ccc aaa tot tgt gac aaa act cac aca tgt cca ccg tcc cca gca cct

<u>HulgGI MTCH3Y405</u> (nucleotide sequence) (SEQ ID NO:\_\_)
gggcageccegagaaccacaggtgtacaccetgcceccatccegggaggagatgaccaaggaacaaggtcagcetgacctgcct
ggtcaaaggcttctatcccagcgacatcgccgtggagtgggagagacaatgggcagcggagaacaactacaagaccacgcctc

HulgG1 MTCH3Y405 (amino acid sequence) (SEQ ID NO:\_\_\_)

GQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPP VLDSDGSFYLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTOKSLSLSPGK

HulgG1 MTCH3A405 (nucleotide sequence) (SEQ ID NO:\_\_\_)

gggeagcccegagaaccacaggtgtacaccctgcccccatcccggaggagatgaccaagaaccaggtcagcctgacctgcct

10 ggtcaaaggcttctatcccagcgacatcgccgtggagtgggagagcaatgggagagcaggggaacaactacaagaccacgcctc

ccgtgctggactccgacggctccttcgccctctatagcaagctcaccgtggacaagagcaggtggcaaggggaacgtcttctc

atgctccgtgatgcatgaggctctgcacaaccactacacgcagaagagcctctccctgtcccgggtaaatga

HuIgG1 MTCH3A405 (amino acid sequence) (SEQ ID NO: \_\_)

15 GQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPP VLDSDGSFALYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSPGK

HuIgG1 MTCH3A407 (nucleotide sequence) (SEQ ID NO:\_\_\_)

<u>HuIgG1 MTCH3A407</u> (amino acid sequence) (SEQ ID NO:\_\_)

25 GQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPP VLDSDGSFFLASKLTVDKSRWOOGNVFSCSVMHEALHNHYTOKSLSLSPGK

HulgG1 MTCH3Y405A407 (nucleotide sequence) (SEQ ID NO: )

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gggcagccccgagaaccacaggtgtacaccctgccccatcccgggaggagatgaccaagaaccaggtcagctgacctgct ggtcaaaggettctatcccagcgacatcgccgtggagtgggagagcaatgggcagcggggagacaactacaagaccacgctc ccgtgctggactccgacggctccttctacctcgccagcaagctcaccgtggacaagagcaggtggcagcaggtggcagcgctttctcattgcccgtgatgcatgagagcgctctccctgtcccgggtaaatga

HulgG1 MTCH3Y405A407 (amino acid sequence) (SEQ ID NO:\_\_)
GQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPP
VLDSDGSFYLASKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSPGK

HulgG1 MTCH3A405A407 (amino acid sequence) (SEQ ID NO:\_\_)
GQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPP
VLDSDGSFALASKLTVDKSRWQQGNVFSCSVMHEALHNHYTOKSLSLSPGK

- 10 2H7 scfv MTH (SSS) WTCH2MTCH3V405 (amino acid sequence) (SEQ ID NO:\_\_)

  MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY

  QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS
  FNPPTFGAGTKLELKDGGGSGGGGGGGGGSQQAYLQQSGAELVRPGASVKMSCK

  ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS

  TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSSDK

  THTSPPSPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV
- THISPPSPAPELIGGESVILPPKEPKDITAMBKIPEVICVVVDVSHEDPEVKENWYV
  DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALFAPIE
  KTISKAKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPPVLDSDGSFYLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS
  20 PGK

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2H7 scFv MTH (SSS) WTCH2MTCH3A405 (nucleotide sequence) (SEQ ID NO:\_\_\_\_ mdfqvqifsfllisasviiargqivlsqspailsaspgekvtmtcrasssvsymhwyqqkpgsspkpwiyapsnlasgvparf sgssgsgtsysltisrveaedaatyycqqwsfnpptfgagtklelkdgggsggggsgsgsgsgaylqqsgaelvrpgasvkmsc kasgytftsymmhwvkqtprqglewigaiypgngdtsynqkfkgkatltvdkssstaymqlssltsedsavyfcarvvyysn sywyfdvwgtgttvtvssdqepkssdkthtsppspapellggpsvflfppkpkdtlmisrtpevtcvvvdvshedpevkfnw yvdgvcvhnaktkpreeqynstyrvvsvltvlhqdwlngkeykckvsnkafpapiektiskakgqprepqvytlppsreemtiknqvsltclvkgfypsdiavewesngqpennykttppvldsdgsfalyskltvdksrwqqgnvfscsvmheallnhytqksl

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gtggtggacgtgagccacgaagaccttgaggtcaagttcaacttgatacgtggacgcgtggaggtgcataatgccaagacaaag ccgegggaggagcagtacaacagcacgtacgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaagga gtacaagtgcaaggtctccaacaaagccctccagccccatcgagaaaacaatctccaaagccaaagggcagcccgagaac cacaggtgtacacctgccccatcccgggaggagtagaccaaggaacaaggacaggtcagctgacctgctggtcaaaggcttctatcc cagegacatcgccgtggagtgggagagcaatgggcagcggagaacaactacaagaccacgcctcccgtgctggactcgac ggctccttcttctctcgccagcaagctcaccgtggacaagagcatgggcagggggaacgacggggaacgtcttctcatgctccgtgatgcatga ggctctgcacaaccactacacgcagaagagctctccctgtcccggggaaatga

2H7 scFv MTH (SSS) WTCH2MTCH3Y405A407 (nucleotide sequence) (SEQ ID

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2H7 scFv MTH (SSS) WTCH2MTCH3Y405A407 (amino acid sequence) (SEQ ID NO: \_\_)

 $\label{thm:model} MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY\\ QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS\\$ 

- 15 FNPPTFGAGTKLELKDGGGSGGGGGGGGGGSQAYLQQSGAELVRPGASVKMSCK
  ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS
  TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSSDK
  THTSPPSPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV
  DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIE
- 20 KTISKAKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPEN NYKTTPPVLDSDGSFYLASKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS PGK

# 2H7 scFv MTH (SSS) WTCH2MTCH3A405A407 (nucleotide sequence) (SEQ ID

25 NO:\_\_\_)

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# 2H7 scFv MTH (SSS) WTCH2MTCH3A405A407 (amino acid sequence) (SEQ ID

20 TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSSDK THTSPPSPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIE KTISKAKGQPREPQVYTLPPSREEMTKNQVSLTCLVKGFYPSDIAVEWESNGQPEN NYKTTPPVLDSDGSFALASKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS

25 PGK

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#### 2H7 scFv MTH (SCC) WTCH2CH3 (nucleotide sequence)

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#### 2H7 scFy MTH (SCC) WTCH2CH3 (amino acid sequence)

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS
FNPPTFGAGTKLELKDGGGSGGGGSGGGGSSQAYLQQSGAELVRPGASVKMSCK
20 ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSSDK
THTCPPCPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWY
VDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPI
EKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPE
25 NNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSL
SPGK

#### 2H7 scFv MTH (CSC) WTCH2CH3 (nucleotide sequence)

aagottgoogocatggatttteaagtgoagattttcagottootgotaateagtgottcagtcataattgocagaggacaaattgitotot cocagtetocagcaatcotgtotgoatotocaggggagaaggtoacaatgacttgoagggocagotcaagtgtaagttacatgoact ggtaccagcagaagcoaggatcotococcaaaccotggatttatgococatccaacctggottotggagtcotgctoctgotgettcagtg

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### 2H7 scFv MTH (CSC) WTCH2CH3 (amino acid sequence)

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS
20 FNPPTFGAGTKLELKDGGGSGGGGSGGGGSQQAYLQQSGAELVRPGASVKMSCK
ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSCDK
THTSPPCPAPELLGGPSVFLFPFKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWY
VDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPI
25 EKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPE
NNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSL
SPGK

#### 2H7 scFv MTH (CCS) WTCH2CH3 (nucleotide sequence)

30 aagettgeegecatggatttteaagtgeagattticagetteetgetaateagtgetteagteataattgeeagaggaeaaattgtietet ceeagteteeageaateetgtetgeateteeaggggagaaggteacaatgaettgeagggeeageteaagtgtaagttaaatgeact

ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg g cagtgggtetgggacctettacteteteaeaateageagagtggaggetgaagatgetgeeacttattactgceageagtggagttttaacccacccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggttggatctggaggaggtg ggageteteaggettatetacageagtetgggetgagetggtgaggeetggggeeteagtgaagatgteetgeaaggettetgge 5 ggtgatacttectacaateagaagtteaagggeaaggeeaeactgaetgtagacaaateeteeageacageetacatgeageteag cagcet gacatet gaag actet geggtet at titet gt geaag ag t g t g tactat ag taactet tact g g tact t e g tactat g tactat gaag actet gaag actet gaag actet g ag g cac actet gaag actet g ag g cac actet g acteagggaccacggtcaccgtctcttctgatcaggagcccaaatcttgtgacaaaactcacacatgtccaccgtccccagcacctgaac tectggggggaccgtcagtcttcctcttcccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcacatgcgtg 10 ccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggagtacaagtgcaaggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaagccaaagggcagccccgagaac cagegacategeegtggagtgggaggagaatgggeageeggagaacaactacaagaccaegeeteeegtgetggaeteegae 15 ggeteettetteetetaeageaageteaeegtggaeaagageaggtggeageaggggaaegtetteteatgeteegtgatgeatga ggetetgeacaaceactacaegeagaagageeteteeetgteteegggtaaatgatetaga

# 2H7 scFv MTH (CCS) WTCH2CH3 (amino acid sequence)

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY

20 QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS
FNPPTFGAGTKLELKDGGGSGGGGGGGGGGSQQAYLQQSGAELVRPGASVKMSCK
ASGYTFTSYMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVVSDQEPKSCDK
THTCPPSPAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWY

25 VDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPI
EKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPE
NNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSL
SPGK

## 30 HulgAHlgA-T4-ORF (nucleotide sequence)

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## HulgAHlgA-T4-ORF (amino acid sequence)

DQPVPSTPPTPSPSTPPTPSPSCCHPRLSLHRPALEDLLLGSEAILTCTLTGLRDASGV
TFTWTPSSGKSAVQGPPDRDLCGCYSVSSVLPGCAEPWNHGKTFTCTAAYPESKT
PLTATLSKSGNTFRPEVHLLPPPSEELALNELVTLTCLARGFSPKDVLVRWLQGSQ
ELPREKYLTWASRQEPSQGTTTFAVTSILRVAAEDWKKGDTFSCMVGHEALPLAF
TOKTIDRLAGKPTHVNVSVVMAEVDADPSN

#### 1D8-IgAH IgA-T4-CD80 (nucleotide sequence)

AA

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#### 10 1D8 scFv IgAH IgA-T4-CD80 (amino acid sequence)

MDFQVQIFSFLLISASVIMSRGVDIVLTQSPTTIAASPGEKVTITCRASSSVSYMYWY QQKSGASPKLWIYDTSKLASGVPNRFSGSGSGTSYSLAINTMETEDAATYYCQQW SSTPLTFGSGTKLEIKRGGGGSGGGGGGGGGGQVQLKEAGPGLVQPTQTLSLTCTV SGFSLTSDGVHWIRQPPGKGLEWMGIIYYDGGTDYNSAIKSRLSISRDTSKSQVFLK

- 15 INSLQTDDTAMYYCARIHFDYWGQGVMVTVSSDQPVPSTPPTPSPSTPPTPSPSCC
  HPRLSLHRPALEDLLLGSEAILTCTLTGLRDASGVTFTWTPSSGKSAVQGPPDRDL
  CGCYSVSSVLPGCAEPWNHGKTFTCTAAYPESKTPLTATLSKSGNTFRPEVHLLPP
  PSEELALNELVTLTCLARGFSPKDVLVRWLQGSQELPREKYLTWASRQEPSQGTTT
  FAVTSILRVAAFDWKKGDTFSCMVGHEALPLAFTQKTIDRLAGKPTHVNVSVVM
- 20 AEVDADPSNNLLPSWAITLISVNGIFVICCLTYCFAPRCRERRNERLRRESVRPV

#### human IgE Fc (CH2-CH3-CH4) ORF (nucleotide sequence)

AA

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#### 5 human IgE Fc (CH2-CH3-CH4) ORF (amino acid sequence)

DHVCSRDFTPPTVKILQSSCDGGGHFPPTIQLLCLVSGYTPGTINITWLEDGQVMDV
DLSTASTTQEGELASTQSELTLSQKHWLSDRTYTCQVTYQGHTFEDSTKKCADSN
PRGVSAYLSRPSPFDLFIRKSPTITCLVVDLAPSKGTVNLTWSRASGKPVNHSTRKE
EKQRNGTLTVTSTLPVGTRDWIEGETYQCRVTHPHLPRALMRSTTKTSGPRAAPE
VYAFATPEWPGSRDKRTLACLIQNFMPEDISVQWLHNEVQLPDARHSTTQPRKTK
GSGFFVFSRLEVIRAEWEQKDEFICRAVHEAASPSQTVQRAVSVNPGKADPS

### 1D8 scFv-human IgE Fc (CH2-CH3-CH4)-CD80 (nucleotide sequence)

aagettatggatttteaagtgeagatttteagetteetgetaateagtgetteagteataatgteeagaggagtegacattgtgeteacte 15 agtotecaacaaccatagetgeateteeaggggagaaggteaceateacetgeegtgeeageteeagtgtaagttacatgtactggt accagcagaagtcaggcgcctcccctaaactctggatttatgacacatccaagctggcttctggagttccaaatcgcttcagtggca gtgggtctgggacctettattetetegeaateaacaceatggagaetgaagatgetgecacttattaetgteageagtggagtagtaet gateteaggtgeagetgaaggaggaggacetggeetggtgeaacegaeacagaceetgteeeteacatgeaetgteietgggtt 20 cacagattataattcagcaattaaatccagactgagcatcagcagggacacctccaagagccaagttttcttaaaaatcaacagtctg caaactgatgacacagccatgtattactgtgccagaatccactttgattactggggccaaggagtcatggtcacagtctcctctgatc acgtetgetecagggaetteaccccgcccaccgtgaagatettacagtegteetgegacggegggggggaetteeccccgaccat 25 ttgtccaccgcctctaccacgcaggaggtgagctggcctccacacaaagcgagctcaccctcagccagaagcactggctgtca gacegeacetacacetgecaggteacetateaaggteacacetttgaggacageaceaagaagtgtgeagatteeaaceegagag gggtgagegeetacetaageeggeeeageeegttegaeetgtteateegeaagtegeeeaegateacetgtetggtggtggaeet ggcacccagcaaggggaccgtgaacctgacctggtcccgggccagtgggaagcctgtgaaccactccaccagaaaggaggag aagcagegcaatggcacgttaaccgtcacgtcaccctgccggtgggcacccgagactggatcgagggggagacctaccagtg 30 cagggtgacccacccccactgcccagggccctcatgcggtccacgaccaagaccagggcccgcgtgctgccccggaagtct atgcgtttgcgacgccggagtggccggggagccgggacaagcgcaccctcgcctgatccagaacttcatgcctgaggac

#### 1D8-scFv-human IgE Fc (CH2-CH3-CH4)-CD80 (amino acid sequence)

MDFQVQIFSFLLISASVIMSRGVDIVLTQSPTTIAASPGEKVTITCRASSSVSYMYWY QQKSGASPKLWIYDTSKLASGVPNRFSGSGSGTSYSLAINTMETEDAATYYCQQW SSTPLIFGSGTKLEIKRGGGGSGGGSGGGGSQQVQLKEAGPGLVQPTQTLSLTCTV SGFSLTSDGVHWIRQPPGKGLEWMGIIYYDGGTDYNSAIKSRLSISRDTSKSQVFLK INSLQTDDTAMYYCARIHFDYWGQGVMVTVSSDHVCSRDFTPFTVKILQSSCDGG GHFPPTIQLLCLVSGYTPGTINITWLEDGQVMDVDLSTASTTQEGELASTQSELTLS QKHWLSDRTYTCQVTYQGHTFEDSTKKCADSNPRGVSAYLSRPSPFDLFIRKSPTI TCLVVDLAPSKGTVNLTWSRASGKPVNHSTRKEEKQRNGTLTVTSTLPVGTRDWI EGETYQCRVTHPHLPRALMRSTTKTSGPRAAPEVYAFATPEWPGSRDKRTLACLI QNFMPEDISVQWLINEVQLPDARHSTTQPRKTKGSGFFVFSRLEVTRAEWEQKDE FICRAVHEAASPSQTVQRAVSVNPGKADPSKLPSWAITLISVNGIFVICCLTYCFAP RCBFRRNPERIRSSVPPV

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### 5B9-IgAH IgA-T4-CD80 (nucleotide sequence)

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#### 5B9-IgAH IgA-T4-CD80 (amino acid sequence)

MRFSAQLLGLLVLWIPGSTADIVMTQAAFSNPVTLGTSASISCRSSKSLLHSNGITY

LYWYLQKPGQSPQLLIYQMSNLASGVPDRFSSSGSGTDFTLRISRVEAEDVGVYYC

AQNLELPLIFGAGTKLELKRGGGSSGGGGSGGGGSSQVQLKQSGPGLVQSSQSLS

ITCTVSGFSLTTYAVHWVRQSPGKGLEWLGVIWSGGITDYNAAFISRLSITKDDSK

SQVFFKMNSLQPNDTAIYYCARNGGDNYPYYYAMDYWGQGTSVTVSSDQPVPST

PPTPSPSTPPTPSPSCCHPRLSLHRPALEDLLLGSEAILTCTLTGLRDASGVTFTWTPS

20 SGKSAVQGPPDRDLCGCYSVSSVLPGCAEPWNHGKTFTCTAAYPESKTPLTATLS

KSGNTFRPEVHLLPPPSEELALNELVTLTCLARGFSPKDVLVRWLQGSQELPREKY

LTWASRQEPSQGTTTFAVTSILRVAAEDWKKGDTFSCMVGHEALPLAFTQKTIDR

LAGKPTHVNVSVVMAEVDADPSNNLLPSWAITLISVNGIFVICCLTYCFAPRCRER

RRNFRI RRFSVRPV

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#### 5B9-scFv-human IgE Fc (CH2-CH3-CH4)-CD80 (nucleotide sequence)

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cgggtggcggcggatcgtcacaggtgcagctgaagcagtcaggacctggcctagtgcagtcctcacagagcctgtccatcacct gcacagtetetggttteteattaactacetatgetgtacactgggttegccagtetecaggaaagggtetggatggetgggtgat atggagtggtggaatcacagactataatgcagctttcatatccagactgagcatcaccaaggacgattccaagagccaagttttcttt aaaatgaacagtctgcaacctaatgacacagccatttattactgtgccagaaatgggggtgataactacccttattactatgctatgga 5 ctactggggtcaaggaacctcagtcaccgtctcctctgatcacgtctgctccagggacttcaccccgcccaccgtgaagatcttacagtcgtcctgcgacggcggcgggcacttcccccgaccatccagctcctgtgcctcgtctctgggtacaccccagggactatcaac ateacetggetggaggaegggeaggteatggaegtggaettgteeaeegeetetaceaegeaggagggtgagetggeeteeaea can age gage teaccet cage caga age act gget gt caga ceg cacet acacet ge cag gt cacet at can gg teaccet tit gage to the category of the categ10 teegeaagtegeeeagateaeetgtetggtggtggaeetggeaeeageaaggggaeegtgaaeetgaeetggteeeggeea gtgggaagcetgtgaaccactccaccagaaaggaggagaagcagcgcaatggcacgttaaccgtcacgtcaccctgccggtg cgaccaagaccageggcccgcgtgctgccccggaagtctatgcgtttgcgacgccggagtggccgggagacaagcgcaccetegcetgatecagaactteatgeetgaggacateteggtgcagtggctgcacaaegaggtgcageteceggaege ccggcacagcacgacgcacgcacgcaagaccaagggctccggcttcttcgtcttcagccgcctggaggtgaccagggccgaat,gggagcagaaagatgagtteatetgeegtgeagteeatgaggeagegageeecteacagacegteeagegageggtgtetgtaa atcccggtaaagcggatccttcgaagctcccatcctgggccattaccttaatctcagtaaatggaatttttgtgatatgctgcctgacct actgctttgccccaagatgcagagagagaaggaggaatgagagttgagaagggaaagtgtacgccctgtataaatcgata

#### 20 5B9-scFv-human IgE Fc (CH2-CH3-CH4)-CD80 (amino acid sequence)

MRFSAOLLGLLVLWIPGSTADIVMTOAAFSNPVTLGTSASISCRSSKSLLHSNGITY LYWYLOKPGOSPOLLIYOMSNLASGVPDRFSSSGSGTDFTLRISRVEAEDVGVYYC AONLELPLTFGAGTKLELKRGGGGGGGGGGGGGGSSOVOLKOSGPGLVOSSOSLS ITCTVSGFSLTTYAVHWVRQSPGKGLEWLGVIWSGGITDYNAAFISRLSITKDDSK 25 SOVFFKMNSLOPNDTAIYYCARNGGDNYPYYYAMDYWGOGTSVTVSSDHVCSR DFTPPTVKILOSSCDGGHFPPTIOLLCLVSGYTPGTINITWLEDGOVMDVDLSTAS TTOEGELASTOSELTLSOKHWLSDRTYTCOVTYOGHTFEDSTKKCADSNPRGVSA YLSRPSPFDLFIRKSPTITCLVVDLAPSKGTVNLTWSRASGKPVNHSTRKEEKORNG TLTVTSTLPVGTRDWIEGETYOCRVTHPHLPRALMRSTTKTSGPRAAPEVYAFATP 30 EWPGSRDKRTLACLIONFMPEDISVOWLHNEVOLPDARHSTTOPRKTKGSGFFVFS

RLEVTRAEWEOKDEFICRAVHEAASPSOTVORAVSVNPGKADPSKLPSWAITLISV NGIFVICCLTYCFAPRCRERRRNERLRRESVRPV

### 2e12-scFv-IgAH IgA-T4-CD80 (nucleotide sequence)

a agettat ggatttt caagt geagatttt cagetteet getaat cagt get teagteat aat gtee ag ag gag te gae at tigt gete acceptation of the properties of the propertiea at ctcc agettettt gget gt et ct aggte ag ag ag ac accate te et ge ag ag ee ag t gaa ag t gt t gaa tat tat gt eac ag t gt tat ge ag ag t gaa ag t gt t gaa tat tat gt eac ag t gt t gaa tat tat gt eac ag t gt t gaa tat tat gt eac ag t gt t gaa tat tat gt eac ag t gt t gaa tat tat gt eac ag t gt t gaa tat tat gt eac ag t gt t gaa tat tat gt eac ag t gt t gaa tat tat gt eac ag t gt t gaa tat tat gt eac ag t gt t gaa tat tat gt eac ag t gt t gaa tat tat gt eac ag t gt t gaa t g5 ta at g cag t g g tacca a cag a a accag ga cag coecca a act cet cat et et g ct g cat ce a acg t a g a at ct g g g g t c c et g cet g cat et a consideration of the state ofaaagtaggaaggtteettggaegtteggtggaggcaecaagetggaaatcaaaeggggtggeggtggetegggeggaggtggg tegggtggeggeggateteaggtgeagetgaaggagteaggacetggeeggtgggegeeeteaeagageetgteeateaeatge accgteteagggtteteattaaccggetatggtgtaaactgggttegecagcetecaggaaagggtetggagtggetgggaatgat 10 atggggtgatggaagcacagactataattcagctctcaaatccagactgagcatcaccaaggacaactccaagagccaagitttcti aaaaatgaacagtetgcaaactgatgacacagccagatactactgtgccagagatggttatagtaactttcattactatgttatggact actgggglcaaggaacetcagtcaccgtctcctcagatcagccagttccctcaactccacctaccccatctccctcaactccaccta cccateteceteatgetgecacecegaetgtcaetgeaecgaeeggeeetegaggaeetgetettaggttcagaagegateete acgtgcacactgaccggcctgagagatgcctcaggtgtcaccttcacctggacgccctcaagtgggaagagcgctgttcaaggac 15 cacetgaeegtgaeetetgtggetgetaeagegtgteeagtgteetgeegggetgtgeegageeatggaaeeatgggaagaeette. A acttgcactgctgcctaccccgagtccaagaccccgctaaccgcaccctctcaaaatccggaaacacattccggcccgaggtcc acctgctgccgccgccgccgcggaggagctggccctgaacgagctggtgacgctgacgtgcctggcacgtggcttcagccccaag .4 gatgtgctggttcgctggtgcaggggtcacaggagctgccccgcgagaagtacctgacttgggcatcccggcaggagcccag 🖟 ccagggcaccaccaccttcgctgtgaccagcatactgcgcgtggcagccgaggactggaagaagggggacaccttctcctgcat 20 gttgtcatggeggaggtggacgggatcettegaacaacetgeteccatcetgggccattacettaatetcagtaaatggaatttttgt gatatgctgcctgacctactgctftgccccaagatgcagagagagaggaggaatgagagagtgagaagggaaagtgtacgccct gtataaatcgatac

### 25 2e12-scFv-IgAH IgA-T4-CD80 (amino acid sequence)

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MDFQVQIFSFLLISASVIMSRGVDIVLTQSPASLAVSLGQRATISCRASESVEYYVTS
LMQWYQQKPGQPPKLLISAASNVESGVPARFSGSGSGTDFSLNIHPVEEDDIAMYF
CQQSRKVPWTFGGGTKLEIKRGGGGSGGGGSGGGGSQVQLKESGPGLVAPSQSLS
ITCTVSGFSLTGYGVNWVRQPPGKGLEWLGMIWGDGSTDYNSALKSRLSITKDNS
KSQVFLKMNSLQTDDTARYYCARDGYSNFHYYVMDYWGQGTSVTVSSDQPVPS
TPPTPSPSTPPTPSPSCCHPRLSLHRPALEDLLLGSEAILTCTLTGLRDASGVTFTWTP
SSGKSAVOGPPDRDLCGCYSVSSVLPGCAEPWNHGKTFTCTAAYPESKTPLTATI.S

KSGNTFRPEVHLLPPPSEELALNELVTLTCLARGFSPKDVLVRWLQGSQELPREKY LTWASRQEPSQGTTFFAVTSILRVAAEDWKKGDTFSCMVGHEALPLAFTQKTIDR LAGKPTHVNVSVVMAEVDADPSNNLLPSWAITLISVNGIFVICCLTYCFAPRCRER RRNERLRRESVRPV

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## 2e12-scFv-human IgE Fc (CH2-CH3-CH4)-CD80 (nucleotide sequence)

aagettatggattticaagtgeagattticagetteetgetaateagtgetteagteataatgteeagaggagtegacattgtgeteacee aatctccagettctttggctgtgtctctaggtcagaggccaccatctcctgcagagccagtgaaagtgttgaatattatgtcacaagtt taatgeagtggtaccaacagaaaccaggacagccacccaaactectcatctctgctgcatccaacgtagaatctggggtccctgcc aggttt agt gg cagt gg gacagact teag ceteaa catecate ctg t gg agg aggat gat at t gcaat gt at tte t gt cagetogggtggegggatet caggtgeagetgaaggagteaggacetggectggtggegeceteacagagectgteeateacatgeaccgtctcagggttctcattaaccggctatggtgtaaactgggttcgccagcctccaggaaagggtctggagtggctgggaatgat atggggtgatggaagcacagactataattcagctctcaaatccagactgagcatcaccaaggacaactccaagagccaagttttctt ... aaaaatgaacagtotgcaaactgatgacacagccagatactactgtgccagagatggttatagtaactttcattactatgttatggact և actggggtcaaggaacctcagtcaccgtctccteagatcacgtctgetccagggacttcaccccgcccaccgtgaagatcttacag \* tegteetgegaeggegggggggaetteeeeegaeeateeageteetgtgeetegtgtaeaeeeeagggaetateaaeat cacctggetggaggaegggcaggtcatggacgtggacttgtceaccgcctctaccacgcaggagggtgagctggcctccacac aaagcgageteacceteagccagaagcactggetgtcagaccgcacctacacctgccaggtcacctatcaaggtcacacctttga ggaeageaceaagaagtgtgeagattceaacecgagagggtgagegetacetaageeggeecagecegttegacetgtteat gtgggaagcctgtgaaccactccaccagaaaggaggagaagcagcgcaatggcacgttaaccgtcacgtcaccctgccggtg cgaccaagaccagcggccgcgtgctgccccggaagtctatgcgtttgcgacgccggagtggccggggagccgggacaagc gcacectegeetgeetgateeagaactteatgeetgaggacateteggtgeagtggetgeacaaegaggtgeageteeeggaege ceggeacageacgacgeageacgeaagaccaagggeteeggettettegtetteageegeetggaggtgaceagggeegaat ateceggtaaageggateettegaageteeeateetgggeeattaeettaateteagtaaatggaatttttgtggatatgetgeetgaeet 

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MDFQVOIFSFLLISASVIMSRGVDIVLTOSPASLAVSLGORATISCRASESVEYYVTS LMQWYQQKPGQPPKLLISAASNVESGVPARFSGSGSGTDFSLNIHPVEEDDIAMYF CQQSRKVPWTFGGGTKLEIKRGGGGSGGGGGGGGGGQVQLKESGPGLVAPSQSLS ITCTVSGFSLTGYGVNWVROPPGKGLEWLGMIWGDGSTDYNSALKSRLSITKDNS 5 KSOVFLKMNSLOTDDTARYYCARDGYSNFHYYVMDYWGOGTSVTVSSDHVCSR DFTPPTVKILQSSCDGGGHFPPTIQLLCLVSGYTPGTINITWLEDGQVMDVDLSTAS TTQEGELASTQSELTLSQKHWLSDRTYTCQVTYQGHTFEDSTKKCADSNPRGVSA YLSRPSPFDLFIRKSPTITCLVVDLAPSKGTVNLTWSRASGKPVNHSTRKEEKORNG TLTVTSTLPVGTRDWIEGETYQCRVTHPHLPRALMRSTTKTSGPRAAPEVYAFATP EWPGSRDKRTLACLIONFMPEDISVOWLHNEVOLPDARHSTTOPRKTKGSGFFVFS RLEVTRAEWEOKDEFICRAVHEAASPSQTVQRAVSVNPGKADPSKLPSWAITLISV NGIEVICCLTYCEAPRCRERRRNERLRRESVRPV

### 500A2 scFv (nucleotide sequence)

15 atgtigtatacateteageteettgggetittactettetggatiteageeteeagaagtgacatagtgetgacteagacteeageeacte tgtctctaattcctggagaaagagtcacaatgacctgtaagaccagtcagaatattggcacaatcttacactggtatcaccaaaaacc aaaggaggeteeaagggeteteateaagtatgettegeagteeatteetgggateeeeteeagatteagtggcagtggtteggaaac agatttcactctcagcatcaataacctggagcctgatgatatcggaatttattactgtcaacaaagtagaagctggcctgtcacgttcgt 20 gctgcagcagtccggttctgaactagggaaacttggggcctcagtgaaactgtcctgcaagacttcaggctacatattcacagatc atcaaaaattccagggcaaggccacactgactgtagataaaatctctagcacagcctacatygaactcagcagcctgacatctgag gattctgccatctattactgtgcaagaaggccggtagcgacgggccatgctatggactactggggtcaggggatccaagttaccgt ctcctctgatc

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#### 500A2 scFv (amino acid sequence)

MLYTSOLLGLLLFWISASRSDIVLTQTPATLSLIPGERVTMTCKTSQNIGTILHWYH QKPKEAPRALIKYASQSIPGIPSRFSGSGSETDFTLSINNLEPDDIGIYYCQQSRSWPV TFGPGTKLEIKRGGGGSGGGGSGGGGSOVKLOOSGSFLGKPGASVKLSCKTSGYIF TDHYISWVKOKPGESLQWIGNVYGGNGGTSYNOKFQGKATLTVDKISSTAYMEL SSLTSEDSAIYYCARRPVATGHAMDYWGOGIOVTVSSD

NT

5' oligo:

Name : IgGWT3

GTTGTTTTCGAAGGATCCGCTTTACCCGGAGACAGGGAGAGGCTCTT

5 NT

3' oligo:

Name : hIgGWT5

GTTGTTAGATCTGGAGCCCAAATCTTGTGACAAAACTCACACATG

NI

10 5' oligo:

Name : FADD5

Sequence

GTTGTGGATCCTTCGAACCCGTTCCTGGTGCTGCTGCACTCGGTGTCG

NT

15 3' oligo:

Name : FADD3

Sequence

GTTGTTATCGATCTCGAGTTATCAGGACGCTTCGGAGGTAGATGCGTC

NT

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## 20 FADD-CSSCFV (nucleotide sequence)

giggatecttegaaccegitectggtgetgetgetgetgetgetagagetgetgetagagetgagetgagetgacgagetgacgageteaagitecta 
igeettegggegegitggeaaagegeaagetggagegegitgeagageggetagacetagactettetectatgetgetggageagaacga 
eetiggagecegggeacaacegageteetgeggagetgetegeeteeetgegegegeagaacgacetgetgeggegettegagaget 
tegaggeggggeggeggeggeggeggegggggagaagaacactggtgegagaatitaacgicatatgtgataatgigggg 
aaagattggagaaggetggetegteageteaagteteagacacaagategaagaacgaacagaagaacgaaceggagaagataceceggeacetg 
acagagegtgigegggagteactgagaaatetggaagaacacagagagagagaacgeaacagggeceacetggigggggete 
teaggteetgecagatgaacetggiggetgacetggtacaagagtitageagagagagagaacagaacagagagtggggeca
tgiccecgatgtaatgaacetggiggetgacetggiacacegaaacaggagtagagaacagaatagagagtggggeca

## 30 FADD-CSSCFV (amino acid sequence)

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VDPSNPFLVLLHSVSSSLSSSELTELKFLCLGRVGKRKLERVQSGLDLFSMLLEQND LEPGHTELLRELLASLRRHDLLRRVDDFEAGAAAGAAPGEEDLCAAFNVICDNVG KDWRRLARQLKVSDTKIDSIEDRYPRNLTERVRESLRIWKNTEKENATVAHLVGA LRSCOMNLVADLVOEVOOARDLONRSGAMSPMSWNSDASTSEAS

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#### HCD28tm5B (nucleotide sequence)

GTTGTGGATCCTCCCTTTTGGGTGCTGGTGGTGGTGGTGTCCTGGCTTGCTAT
AGCTTG

#### 10 HCD28tm3S (nucleotide sequence)

 $\label{eq:condition} \textbf{GTTGTTCGAACCCAGAAAATAAAAAGGCCACTGTTACTAGCAAGCTATAGCAAGCCAG}$  AAGCCAG

## HCD28tm5' (nucleotide sequence)

15 GTTGTGGATCCTCCCTTTTGGGTGCTGGTGGT

#### HCD28tm3' (nucleotide sequence)

GTTGTTTCGAACCCAGAAAATAATAAAGGCCAC

### 20 HCD80tm5' (nucleotide sequence)

GTTGTGGATCCTCCTGCTCCCATCCTGG

#### HCD80tm3' (nucleotide sequence)

25 GTTGTTTCGAACGGCAAAGCAGTAGGTCAGGC

#### MFADD5BB (nucleotide sequence)

GTTGTGGATCCTTCGAACCCATTCCTGGTGCTGCTGCACTCGCTG

#### MFADD3XC (nucleotide sequence)

#### GTTGTTATCGATCTCGAGTCAGGGTGTTTCTGAGGAAGACAC

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#### Murine FADD (amino acid sequence)

VDPSNMDPFLVLLHSLSGSLSGNDLMELKFLCRERVSKRKLERVQSGLDLFTVLLE QNDLERGHTGLLRELLASLRRHDLLQRLDDFEAGTATAAPPGEADLQVAFDIVCD NVGRDWKRLARELKVSEAKMDGIEEKYPRSLSERVRESLKVWKNAEKKNASVA GLVKALRTCRLNLVADLVEEAOESVSKSENMSPVLRDSTVSSSETP

#### MCASP3-5 (nucleotide sequence)

GTTGTGGATCCTTCGAACATGGAGAACAACAAAACCTCAGTGGATTCA

### 25 MCASP3-3 (nucleotide sequence)

GTTGTTATCGATCTCGAGCTAGTGATAAAAGTACAGTTCTTTCGT

#### MCASP8-5 (nucleotide sequence)

GTTGTTTCGAACATGGATTTCCAGAGTTGTCTTTATGCTATTGCTG

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#### MCASP8-3 (nucleotide sequence)

GTTGTTATCGATCTCGAGTCATTAGGGAGGGAAGAAGAGCTTCTTCCG

#### 5 hcasp3-5(nucleotide sequence)

GTTGTGGATCCTTCGAACATGGAGAACACTGAAAACTCAGTGGAT

#### hcasp3-3 (nucleotide sequence)

GTTGTTATCGATCTCGAGTTAGTGATAAAAATAGAGTTCTTTTGTGAG

10

## hcasp8-5 (nucleotide sequence)

GTTGTGGATCCTTCGAACATGGACTTCAGCAGAAATCTTTATGAT

### <u>hcasp8-3</u> (nucleotide sequence)

15 GTTGTTATCGATGCATGCTCAATCAGAAGGGAAGACAAGTTTTTTCT

#### 1. 2H7 scFv with alternative VHL11 mutations:

Nucleotide sequence

- Amino acid sequence
  MDFQVQIPSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
  QQKPGSSPKPWIYAPSNLASGVPARFSGGGGTSYSLTISRVEAEDAATYYCQQWS
  FNPFTFGGGTKLELKDGGGSGGGGGGGGGSQAVLOOSGAE (one of the followine:
- 35 S, T, D, E, Q, N, R, K, H)
  VRPGASVKMSCKASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFK

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 $\begin{array}{l} {\sf GKATLTVDKSSSTAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVT}\\ {\sf VSSDO} \end{array}$ 

### 2. VHL11 deletion

5 Nucleotide sequence:

### Amino acid sequence:

MDFQVQIFSFLLISASVILARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY QQKPGSSPKPWIYJ APSNLASGVPARFSGSGSTSYSLTISRVEAEDAATYYCQQWS 20 FNPPTFGAGTKLELKDGGGSGGGSGGGGSSQAYLQQSGAEVRPGASVKMSCKA SGYTTTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSST AYMOLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVYSSD

#### 3. 2H7 VL L106 with alternative mutations

Nucleotide sequence:

Amino acid sequence:

MDFQVQIFSFLLISASVILARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS FNPPTFGAGTKLE (S, T, R, K, H, Q, N, D, E, and non-natural derivatives of these amino acids at position 100/KDGGGSGGGGSGGGGSS

#### 4. VL L106 deletion

40 Nucleotide sequence:

A gett geogreat ggatttte agt geagatttte agette og taket agt gette agt att att geografiske til te te en gette og generatiet et generatiet et generatiet et generatiet et generatiet genera

45 tttaacccaccacgttcggtgctgggaccaagctggaganaagatggcggtggctcgggcggtggtggatctggaggaggtgg gagctc

Amino acid sequence:

15

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40

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MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS FNPPTFGAGTKLEKDGGGSGGGGSGGGGSS

### 5. IgE CH3 CH4

Nucleotide sequence:

Amino acid sequence:

SNPRGVSAYLSRPSPFDLFIRKSPTITCLVVDLAPSKGTVNLTWSRASGKPVNHSTR KEEKQRNGTLTVTSTLPVGTRDWIEGETYQCRVTHPHLPRALMRSTTKTSGPRAA PEVYAFATPEWPGSRDKRTLACLIQNFMPEDIS VQWLHNEVQLPDARHSTTQPRK TKGSGFFVFSRLEVTRAEWBOKDEFICRAVHEAASPSOTVORAVSVNPGK

# 6. hIgG1H/IgE WCH3 WCH4

Nucleotide sequence:

Amino acid sequence:

35 DQEPKSSDKTHTSPPSPASNPRGVSAYLSRPSPFDLFIRKSPTITCLVVDLAPSKGTV NLTWSRASGKPVNHSTRKEEK QRNGTLTVTSTLPVGTRDVWEGETY QCRYTHPHL PRALMRSTTKTSGPRAAPEVYAFATPEWPGSRDKRTLACLIQNFMPEDISVQWLH NEVQLPDARHSTTQPRKTKGSGFFVFSRLEVTRAEWEQKDEFICRAVHEAASPSQT VQRAVSVNPGK

### 7. IgE WCH2 WCH3 WCH4

Nucleotide sequence:

WO 2005/017148 PCT/US2003/041600

gaggacaltotegtgoattgottgoacaacgagtgoagotocoggaogocoggoacgacacgaogocogococgoagoco aagggotocoggottottogtottoagocgoctggagtgaacagggocgaatgggagaagaaagatggattaattgocgtgoag tocatgaggoagogagocotoacagacogtocagogagoggtgtotgtaattocoggtaattgataattaga

5 Amino acid sequence:

DHVCSRDFTPPTVKILQSSCDGGGHPPPTIGLICLVSGYTPGTINITWLEDGQVMDV DLSTASTTQEGELASTQSELTLSQKHWLSDRTYTCQVTYQGHTFEDSTKKCADSN PRGVSAYI.SRPSPFDLFIRKSPTITCLVVDLAPSKGTVNLTWSRASGKPVNHSTRKE EKQRNGTLTVTSTLPVGTRDWIEGETYQCRVTHPHLPRALMRSTTKTSGPRAAPE VYAFATPEWPGSRDKRTLACLIQNFMPEDISVQWLHNEVQLPDARHSTTQPRKTK GSGFFVFSRLEVTRABWEQKDEFICRAVHBAASFSOTVQRAVSVNPGK

# 8. hIgG1H/IgE CH3 CH4 (ORF)

Nucleotide sequence:

25 Amino acid sequence:

AMILIO acus sequence:
DQEPKSSDKTHTSPSPASNPRGVSAYLSRPSPFDLFIRKSPTITCLVVDLAPSKGTV
NLTWSRASGKPVNHSTRKEEKQRNGTLTVTSTLPVGTRDWIEGETY QCRVTHEHL
PRALMRSTTKTSGPRAPEVY AF ATPEWPGSRDKRTLACLIQNFMPEDISVQWLH
NEVQLPDARHSTTQPRKTKGSGFFVFSRLEVTRAEWEQKDEFICRAVHEAASPSQT

30 VQRAVSVNPGKSGSFE

# 9. 2H7 VHL11S scFv hIgG1(SSS-S)H hIgE WCH3 WCH4

Nucleotide sequence:

- aagettgeegecatggattiteaagtgeagattiteagetteetgetaateagtgetteagteataattgeeagaggaeaaattgttetet 35 cccagtctccagcaatcctgtctgcatctccaggggagaaggtcacaatgacttgcagggccagctcaagtgtaagttacatgcact ggtaccagcagaagccaggatcetcccccaaaccctggatttatgccccatccaacctggcttctggagtcctgctcgcttcagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtgggageteteaggettatetacageagtetggggetgagteggtgaggeetggggeeteagtgaagatgteetgeaaggettetgge tacacatttaccagttacaatatgcactgggtaaagcagacactagacagggcetggaatggattggagctatttatccaggaaat cagcetgacatetgaagactetgeggtetatttetgtgcaagagtggtgtactatagtaactettactggtacttegatgtetggggcac agggaccacggtcaccgtctttctgatcaggagcccaaatcttctgacaaaactcacacatccccaccgtcctcagcatccaacc cgagagggtgagegeetacetaageeggeeeageeegttegacetgtteateegeaagtegeeeacgaleacetgtetggtggt 45 ggacetggeacecagcaaggggacegtgaacetgacetggtecegggcagtgggaagcetgtgaaccaetceaccagaaag gaggagaagcagcagcaatggcacgttaaccgtcaccctgccggtgggcacccgagactggatcgaggggggagacct accagtgeagggtgacceaceceacetgeceagggccctcatgeggtccacgaccaagaccaggcccgcgtgctgcccc
- ggaagtotatgogtttigogacgcoggaagtggcogggaacaagcgcaccotcgcotgactcagaacttatgcc tagaggacatdcggtgcagtggctgcacaacgaaggtgcagctcocggaagccoggaagcaagcaagcaagcacgcaagcacg agggctcoggattottegtottcagccgcotggaagtgaccagggcogaatggagaagaagaatgattacttgccgtgcag tccafgaggcagcgagcocotcacagacogtccagcgacggtgttgtaaatcocggtaaatgataatctaga

Amino acid sequence:

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY OOKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCOOWS FNPTFGAGTKLELKDGGGSGGGGSGGGSSOAYLOOSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKOTPROGLEWIGAIYPGNGDTSYNOKFKGKATLTVDKSSS TAYMOLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDOEPKSSDK THTSPPSSASNPRGVSAYLSRPSPFDLFIRKSPTITCLVVDLAPSKGTVNLTWSRASG KPVNHSTRKEEKORNGTLTVTSTLPVGTRDWIEGETYOCRVTHPHLPRALMRSTT KTSGPRAAPEVYAFATPEWPGSRDKRTLACLIONFMPEDISVOWLHNEVOLPDAR 10 HSTTQPRKTKGSGFFVFSRLEVTRAEWEQKDEFICRAVHEAASPSQTVQRAVSVNP GK

#### 10. 2H7 VHL11S scFv hIgG1(SSS-P)H hIgE WCH3 WCH4

Nucleotide sequence:

15

2.0

30

35

45

aagettgeegeeatggatttte aagtgeagatttte agetteetgetaate agtgette agteataattgee agaggaeaa attgttetetcccagtetecageaatectgtetgeatetecaggggagaaggteaeaatgaettgeagggeeageteaagtgtaagttaeatgeaet ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtcctgctcgcttcagtg g cagtgggtetgggacctettaeteteteaeaateageagagtggaggetgaagatgetgeeaettattaetgeeageagtggagttttaacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggtctggaggaggtg ggageteteaggettatetacageagtetggggetgagteggtgaggeetggggeeteagtgaagatgteetgeaaggettetgge tacacatttaccagttacaatatgcactgggtaaagcagcagcacctagacagggcctggaatggattggagctatttatccaggaaat agggaccaeggteacegtetettetgatcaggageceaaatettetgacaaaaeteacacatececaeegteeceageatecaaee cgagagggtgagcgcctacctaagccggcccagcccgttcgacctgttcatccgcaagtcgccacgatcacctgtctggtggt accagtg cagggtg acceaecceecctg cccaggg cccteatg cggtccacgaccaugaccag cggcccgcgtgctgccccggaagtetatgegtttgegaegeeggagtggeeggggageegggaeaagegeaceetegeetgetgateeagaaetteatgee tgaggacatctcggtgcagtggctgcacaacgaggtgcagctcccggacgcccggcacagcacgacgcagccccgcaagacc aagggeteeggettettegtetteageegeetggaggtgaceagggeegaatgggagcagaaagatgagtteatetgeegtgeag tecatgaggcagcgagcccctcacagaccgtccagcgagcggtgtctgtaaatcccggtaaatgataatctaga

Amino acid sequence: MDFOVOIFSFLLISASVIIARGOIVLSOSPAILSASPGEKVTMTCRASSSVSYMHWY QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS FNPPTFGAGTKLELKDGGGSGGGGGGGGSSOAYLOOSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKOTPROGLEWIGAIYPGNGDTSYNOKFKGKATLTVDKSSS TAYMOLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDOEPKSSDK THTSPPSPASNPRGVSAYLSRPSPFDLFIRKSPTITCLVVDLAPSKGTVNLTWSRASG KPVNHSTRKEEKORNGTLTVTSTLPVGTRDWIEGETYOCRVTHPHLPRALMRSTT KTSGPRAAPEVYAFATPEWPGSRDKRTLACLIQNFMPEDISVQWLHNEVQLPDAR HSTTOPRKTKGSGFFVFSRLEVTRAEWEOKDEFICRAVHEAASPSOTVORAVSVNP GK

### 10. 2H7 VL L106S

a agett g ccgccat ggattit caagt g cagattit cagett cctgct a at cagtget teagt cata at tgccag aggaca a at tgt tetet50 ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccttgctcgcttcagtg

# 5 Amino acid sequence:

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS FNPPTFGAGTKLESKDOGGSGGGGSGGGGSS

### 10 11. 2H7 VL L106S scEv

Nucleotide sequence:

### Amino acid sequence:

MDFQVQIFSFLLISASVILARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
QQKFGSSPKPWIYAPSNLASGVPARFSGSGSGSTSYSLTISRVEAEDAATYVCQQWS
FNPPTFGAGTKLESKDGGSGGGGGGGGSGGAYLQQSGAELVRPGASVKMSCK
ASGYIFTISYNMHWVKQPPRQGLEWIGAIYPGNODTSYNQKFKGKATLTVDKSSS
TAYMOLSSLTSEDSAVYFCARVYYSNSYWYPDVWGTGTTVTVSSOD

### 30 12. 2H7 scFv VL L106S VHL11S scFv

acagggaccacggtcaccgtctcttctgatcag

Nucleotide sequence:

35

40

Amino acid sequence:

MDFQVQIFSFI.LISASVILARGQIVI.SQSPAILSASPGEKVTMTCRASSSVSYMHWY
QQKPGSSPKPWIY.PASNLASGVPARFSGSGGTSYSLTISRVPAEDAATATYCQQWS
FNPPTFGAGTKLESKDGGGSGGGGSGGGGSSQAYLQQSGAESVRPGASVKMSCK
ASGYTFTSYMHWVKQTPRQGLEWIGATYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYPCARVYYSNSYWYFDVWGTGTTVTVVSSOT

50 10. Human IgD hinge linker with attached restriction sites Nucleotide:

PCT/US2003/041600

gtggatccaggttcgaagtctccaaaggcacaggcctcctccgtgcccactgcacaaccccaagcagagggcagcctcgccaa caagaagagagagacaaagaccggtgcagtcgacg

Amino acid:

VDPGSKSPKAQASSVPTAQPQAEGSLAKATTAPATTRNTGRGGEEKKKEKEKEEQ EERETKTGAVD

### Sequence of Native IgD hinge domain:

10 (includes a cysteine residue-we truncated the hinge prior to that residue for these constructs:)

Nucleotide:

gagtetecaaaggeacaggeeteeteegtgeecactgeacaaceecaageagagggeageetegecaaggeaaceacageee gacaaagacaccagagtgtccgagccacacccagcctcttggcgtctacctgctaacccct

Amino acid sequence: ESPKAQASSVPTAQPQAEGSLAKATTAPATTRNTGRGGEEKKKEKEKEEQEERET KTPECPSHTOPLGVYLLTP

20

15

# 12. 2H7 VH L11S

Nucleotide sequence:

eagget late tacage agtet gggget gag teggt gag geet eagt gaa gat gteet geaa gget tet gg tacae at the contract of the contract of the contract grant gaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaatggtgatact 25 tectacaatcagaagttcaagggcaaggccacactgactgtagacaaatcctccagcacagcctacatgcagctcagcagcctga aeggteacegtetettet

Amino acid sequence:

30 QAYLQQSGAESVRPGASVKMSCKASGYTFTSYNMHWVKQTPRQGLEWIGAIYPG NGDTSYNOKFKGKATLTVDKSSSTAYMOLSSLTSEDSAVYFCARVVYYSNSYWY FDVWGTGTTVTVSS

# 13. 2H7 VH L11S scFv

35 Nucleotide sequence:

> augettgeegecatggatttteaagtgeagatttteagetteetgetaateagtgetteagteataattgeeagaggaeaaattgttetet cccagtctccagcaatcctgtctgcatctccaggggagaaggtcacaatgacttgcagggccagctcaagtgtaagttacatgcact gcagtgggtctgggacctcttactctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacceaccacgtteggtgetgggaccaagetggagetgaaagatggeggtggetegggeggtggtggatetggaggaggtg

- 40 ggageteteaggettatetacageagtetggggetgagteggtgaggeetggggeeteagtgaagatgteetgeaaggettetgge tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat cagcet gacatet gaag actet geggtet att tet gt geaag agt ggt gt act at agt aactet taet gg taet te gat get extended a tender of the control of45 agggaccacggtcaccgtctcttctgatcag

Amino acid sequence:

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY QOKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCOOWS

PCT/US2003/041600

ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS TAYMOLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDO

# 14. 2H7 scFv VH L11S hIgG1 (CSC-S)H WCH2 WCH3

5 Nucleotide sequence:

aagettgccgccatggattttcaagtgcagattticagettectgctaatcagtgettcagtcataattgccagaggacaaattgttetet cccagtetecageaateetgtetgeatetecaggggagaaggteacaatgaettgeagggeeageteaagtgtaagttacatgeaet ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg geagtgggtetgggacetettacteteteacaateageagagtggaggetgaagatgetgeeacttattactgeeageagtggagttt 10 taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggtcgggcggtggtggatctggaggaggtg ggagctctcaggcttatctacagcagtctggggctgagtctgtgaggcctggggcctcagtgaagatgtcctgcaaggcttctggct acacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaatg agcetgacatetgaagactetgeggtetattietgtgeaagagtggtgtactatagtaactettactggtacttegatgtetggggeaca 15 gggaccaeggteacegtetettetgateaggageceaaatettgtgacaaaaeteacacateteeacegtgeteageacetgaacte ctgggtggaccgtcagtcttcctcttcccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcacatgcgtggt ggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagc cgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggag ta caagtg caagg to to caa caa ag coctoc cag cocccat c gagaa aa caato to caa ag gcaa ag gcaag cocc gagaa coccat caa ag gcaa ag gcaag gcaa20 agcgacatcgccgtggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacg geteettetteetetaeageaageteaeegtggacaagageaggtggeageaggggaaegtetteteatgeteegtgatgeatgag getetgeaeaaceaetacaegeagaagageeteteeetgteteegggtaaatgatetaga

25 Amino acid sequence:

MDFQVQIES\*LLISASVILARGQIVLSQSPAILSASPGBEKVTMTCRASSSVSYMHWY
QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS
FNPPTFGAGTKLELKDGGGSGGGSGGGGSGGAYLQQSGAESVRPGASVKMSCK
ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVYYSNSYWYFDVWGTGTTVTVSSDQEPKSCDK
THTSPPCSAPBLLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWV
VDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNKGEYKCKVSNKALPAPI
EKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNQOPE
NNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSL
SFGK

# 15. 2H7 scFv VH L11S IgE WCH2 WCH3 WCH4

Nucleotide sequence:

aagettgeegeeatggatttteaagtgeagatttteagetteetgetaateagtgetteagteataattgeeagaggaeaaattgttetet 40 cccagtctccagcaatcctgtctgcatctccaggggagaaggtcacaatgacttgcagggccagctcaagtgtaagttacatgcact ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggageteteaggettatetaeageagtetggggetgagtetgtgaggeetgggggeeteagtgaagatgteetgeaaggettetgget 45 acacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcetggaatggattggagctatttatccaggaaatg gtgataetteetaeaateagaagtteaagggeaaggeeacaetgaetgtagaeaaateeteeageacageetacatgeageteage agectgacatetgaagactetgeggtetattietgtgeaagagtggtgtactatagtaactettaetggtacttegatgtetggggeaea gggaccacggtcaccgtctcttctgatcacgtctgctccagggacttcaccccgcccaccgtgaagatcttacagtcgtcctgcgac ggeggegggactteeccegaccatecageteetgtgeetegtetetgggtacaccecagggactateaacateacetggetgg 50 aggacggcaggtcatggacgtggacttgtccaccgcctctaccacgcaggagggtgagctggcctccacacaaagcgagctc acceteagecagaageaetggetgteagacegeacetacacetgecaggteacetateaaggteacacetttgaggacageacea

10 Amino acid sequence:

MDFQVQIESFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS
FNPPTFGAGTKLELKDGGGSGGGGGGGGGSGGAGGAEVRPGASVKMSGK
15 ASGYTFTSYNMEWVKQTPRQGLEWIGATYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDHVCSRDFT
PPTVKILQSSCDGGHPPPTIQLLCLVSGYTPGTINITWLEDGQVMDVDLSTASTTQ
EGELASTQSELTLSQKHWLSDRTYTCQVTYQGHTFEDSTKKCADSNPRGVSAYLS
RSSFDLFIRKSPTTTCLVVDLAPSKGTVNLTWSR ASGKPVNHSTRKEEKQENGTLT.
20 VISTLPVGTRDWIEGETYQCRVTHPHLPRALMRSTTKTSGPRAAPEVYAFATPEW
PGSRDKRTLACLIQNFMPEDISVQWLHNEVQLPDARHSTTQPRKTKGSGFFVFSRL
EVTRAEWEGKDEFICRAVHEAASPSOTVORAVSVNPGK

# 16. 2H7 scFv VH L11S mIgE WCH2 WCH3 WCH4

Nucleotide sequence:

25

a agett geogecat gg att tica agt ge agatt tica gette et get a at caging et te agt cat a att georga gas a aut gette et gette agt cat a te georga gas a aut gette et gette agt cat a te georga gas a aut gette et gette et gette agt cat a te gette et gette etcccagtetecageaateetgtetgeateteeaggggagaaggteaeaatgaettgeagggeeageteaagtgtaagttacatgeaet ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggettctggagtccctgctcgcttcagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt 30 taacccaccacgtteggtgetgggaccaagctggagetgaaagatggeggtggtegggtggtggtggatetggaggaggtg ggageteteaggettatetacageagtetggggetgagtetgtgaggeetggggeetcagtgaagatgteetgeaaggettetgget acacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcetggaatggattggagctatttatccaggaaatg agcctg a catctg a agactctg cggtct atttctg tg caa gagtg gtg tactatag taactcttactg gtacttc gatgtctg ggg caca35 gggaccacggtcaccgtctcttctgatcacgttcgacctgtcaacatcactgagcccaccttggagctactccattcatcctgcgacc gggagataactgatacacttgcacaaactgttctaatcaaggaggaaggcaaactagcctctacctgcagtaaactcaacatcactg agcagcaatggatgtetgaaagcacettcacetgcaaggtcaceteccaaggegtagactatttggeccacacteggagatgceca gateatgagecaeggggtgtgattacetacetgateceaeceagececetggacetgtateaaaaeggtgeteeeaagettaeetgt 40 ctggtggtggacctggaaagcgagaagaatgtcaatgtgacglggaaccaagagaagaagacttcagtctcagcatcccagtggt acactaagcaccacaataacgccacaactagtatcacctccatcetgcctgtagttgccaaggactggattgaaggctacggctatc agtgeatagtggaccaccctgattttcccaagcccattgtgcgttccatcaccaagaccccaggccagcgctcagcccccgaggtatatgtgttcccaccaccagaggaggaggaggagagacaaacgcacatcacctgtttgatccagaacttcttcctgaggatatetet gtgcagtggctgggggatggcaaactgatetcaaacagecagcacagtaccacaacacccctgaaatccaatggctccaatcaa 45 ggettetteatetteagtegeetagaggtegeeaagaeaetetggaeaeagagaaaaeagtteaeetgeeaagtgateeatgagge actteagaaacceaggaaactggagaaaacaatatecacaagcettggtaacaccteceteegtecatectagtaatctagag

Amino acid sequence:

MDFQVQIRSFILISASVIIARGQIVI.SQSPAILSASPGEKVTMTCRASSSVSYMHWY
QQKPGSSPKPWIYAPSNLASGVPARFSGGSGTSYSLTISRVEAEDAATYYCQQWS
FNPPTFGAGTKLELKDGGGSGGGGGGGGGGSQAYLQQSGAESVRPGASVKMSCK

PCT/US2003/041600

ASGYTFTSYNMHWVKQTPRQGLEWIGATPGNGDTSYNQKEKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDHVRPVNIT
EPTLELLHSSCDPNAFHSTIQLYCFIYGHILNDVSVSWLMDDREITDTLAQTVLIKE
EGKLASTCSKLNITEQQWMSESTFTCKVTSQGVDVLAHTRRCPDHEPRGVTTYLD
PSPILDLYQNGAPKLTGLVVDLESEKNVNTVHNQKKTSVSASQWYTKHHNNATT
SITSILPVVAKDWIEGYGYQCIVDHPDFPKPIVRSITKTPGQRSAPFVYVFPPPEEISE
DKRTLTCLIQNFFPEDISVQULGDGKLISNSQHSTTTPLKSNGSNQGFFIFSRLEVAK
TLWTORKOFTCOVHEALOKPRKLEFTISTSLGNTSI RPS

# 10 17. 2H7 scFv VH L11S hIgA WH WCH2 T4CH3 Nucleotide sequence:

cccag to tecag caate ct g to tecag g g agaa g g to a caat g act t g cag g c cag ctca a g t g taa g t tacat g cact a cat g cact g cag g cag ctca a g t g taa g t tacat g cact g cag g cag ctca a g t g taa g t tacat g cact g cac g ctca a g t g t a g t cac g cac g ctca g cac g cac g ctca g cac g cac g ctca g cac g ca ${\tt ggtaccagcagaagccaggatectcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg}$ 15 gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggagctetcaggcttatctacagcagtctggggctgagtctgtgaggcctggggcctcagtgaagatgtcctgcaaggcttctggct acacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaatg 20 ageetgacatetgaagactetgeggtetatttetgtgeaagagtggtgtactatagtaactettactggtacttegatgtetggggeaca gggaccacggtcaccgtctcttctgatcagccagttccctcaactccacctaccccatctccctcaactccacctaccccatctccct catgctgccaccccgactgtcactgcaccgaccggccctcgaggacctgctcttaggttcagaagcgatcctcacgtgcacactg accggcctgagagatgcctcaggtgtcaccttcacctggacgccctcaagtgggaagagcgctgttcaaggaccacctgaccgtg acctctgtggctgctacagcgtgtccagigtcctgccgggctgtgccgagccatggaaccatgggaagaccttcacttgcactgct 25 geetaeeeegagteeaagaeeeegetaaeegeeaceeteteaaaateeggaaaeacatteeggeeegaggteeaeetgetgeeg ccgccgtcggaggagctggccctgaacgagctggtgacgctgacgtgcctggcacgtggcttcagccccaaggatgtgctggtt ccaccttcgctgtgaccagcatactgcgcgtggcagccgaggactggaagaagggggacaccttctcctgcatggtgggccacg 30 gaggtggactgataatctaga

Amino acid sequence:

MDFQVQIESİLLISASVILARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMIRVY
QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS
FNPFTFGAGTKLELKDGGGSGGGSGGGGSSQAYLQQSGAESVRPGASVKMSCK
ASGYFTISYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSBDSAVYFCARVYYSNSVWFDVWGTGTTTVTVSSDQPVPSTPPT
PSPSTPPTPSPSCCHPRI.SLHRPALEDLI.GSEAILTCTLTGLRDASGVTFTWTPSSG
KSAVQGPPDRDLCGCYSVSSVLPFGCAEPWNHGKTFTCTAAYPESKTPLTATLSKS
GNTFRPEVHLLPPPSEELALNELVTLTCLARGFSPKDVLVRWLQGSQETPEEYLT
WASRQEPSQGTTTFAVTSILRVAAEDWKKGDTFSCMVGHEALPLAFTQKTIDRLA
GKPTTIVNYSVVMAEVD

# 18. 2H7 scFv VH L11S mIgA WH WCH2 T4 CH3

45 Nucleotide sequence:

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a cacatttac cagttaca at at gcactgggta a ag cagaca cct ag a cagggcctgga at tgga gct at tt at ccagga a at gcactgga at the content of the contgtgatacttectae a at caga agtte a aggge a aggee a cact gact gtaga ca a at cete cage acage ctae at geage teage to a geage acage consistent of the contract ofgeggccagctcttgaggacctgctcctgggttcagatgccagcatcacatgtactctgaatggcctgagagatcctgagggagctggtetgeetggetgtgetgagegetggaacagtggegeatcattcaagtgcacagttacccatcctgagtetgacacettaactgge acaattgccaaagtcacagtgaacacettcccaccccaggtccacetgctaccgccgccgtcggaggagctggccctgaatgag ctcgtgtccctgacatgcctggtgcgagctttcaaccctaaagaagtgctggtgcgatggctgcatggaaatgaggagctgtcccc gctgaaatctggaaacagggtgaccagtactcctgcatggtgggccacgaggccttgcccatgaacttcacccagaagaccatcg accgtctgtcgggtaaacccaccaatgtcagcgtgtctgtgatcatgtcagagggagattgataatctagat

### Amino acid sequence:

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY 15 QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS FNPPTFGAGTKLELKDGGGSGGGGSGGGSSQAYLQQSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKOTPROGLEWIGAIYPGNGDTSYNOKFKGKATLTVDKSSS TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDHICSPPTTP PPPSCOPSLSLORPALEDLLLGSDASITCTLNGLRDPEGAVFTWEPSTGKDAVOKK AVONSCGCYSVSSVLPGCAERWNSGASFKCTVTHPESDTLTGTIAKVTVNTFPPOV HLLPPPSEELALNELVSLTCLVRAFNPKEVLVRWLHGNEELSPESYLVFEPLKEPGE GATTYLVTSVLRVSAEIWKOGDOYSCMVGHEALPMNFTOKTIDRLSGKPTNVSVS

A. mIgA WCH2 T4CH3

# Nucleotide sequence:

VIMSEGD

Gttgttgatcacatctgttctcctcctactactcctcctccaccttcctgccageccagcctgtcactgcagegggccagctcttgagga cctgctcctgggttcagatgccagcatcacatgtactctgaatggcctgagagatcctgagggagctgtcttcacctgggagccctc 30 agegetggaacagtggegeateatteaagtgeacagttacecateetgagtetgacaecttaactggeacaattgeeaaagteaca gtgaacacetteceacccaggtceacctgetaccgccgccgtcggaggagctggccctgaatgagctcgtgtccctgacatgcc tggtgegagettteaaccetaaagaagtgetggtgegatggetgeatggaaatgaggagetgteeccagaaagetacetagtgtttg agecectaaaggagecaggegaggagecaceacetacetggtgacaagegtgttgegtgtateagetgaaatetggaaacagg 35 gtgaccagtactcctgcatggtgggccacgaggccttgcccatgaacttcacccagaagaccatcgaccgtctgtcgggtaaacc caccaatgtcagcgtgtctgtgatcatgtcagagggagattgataatctagat

### Amino acid sequence:

DHICSPPTTPPPPSCOPSLSLORPALEDLLLGSDASITCTLNGLRDPEGAVFTWEPST 40 GKDAVQKKAVQNSCGCYSVSSVLPGCAERWNSGASFKCTVTHPESDTLTGTIAKV TVNTFPPOVHLLPPPSEELALNELVSLTCLVRAFNPKEVLVRWLHGNEELSPESYL VFEPLKEPGEGATTYLVTSVLRVSAEIWKOGDOYSCMVGHEALPMNFTOKTIDRL SGKPTNVSVSVIMSEGD

#### 45 K322S CH2 region 20.

Nucleotide sequence:

cetgaaeteetggggggaeegteagtetteetetteececcaaaacecaaggaeacectcatgateteeeggaeecetgaggteae gacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatg geaaggagtacaagtgeteggtetecaacaaageceteceagececeategagaaaacaatetecaaagecaaa

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Amino acid sequence:

PELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEVHN AKTKPREEOYNSTYRVVSVLTVLHODWLNGKEYKCSVSNKALPAPIEKTISKAK

#### 21. K322S CH2 WCH3

Nucleotide sequence:

cetgaaeteetggggggaeegteagtetteetetteeceecaaaaeceaaggaeaeceteatgateteeeggaeeeetgaggteae atgcgtggtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaa 10 gacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatg gcaaggagtacaagtgctcggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaagccaaagggcagccc cttctatcccagcgacatcgccgtggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctgg actccgacggctccttcttcctctacagcaagctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtg 15 atgcatgaggetetgeacaaccactacacgcagaagageeteteeetgteteegggtaaatgatetaga

Amino acid sequence:

PELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEVHN AKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCSVSNKALPAPIEKTISKAKG OPREPOVYTLPPSRDELTKNOVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPPV LDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSPGK

#### K322L CH2 WCH3

Nucleotide sequence:

25 tgatcaggageccaaatcttctgacaaaactcacacatccccacegtectcagcacctgaactcctggggggaccgtcagtcttect cttcccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcacatgcgtggtggtggacgtgagccacgaaga geacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggagtacaagtgcctggtctccaacaa ageceteccageccccategagaaaacaatetecaaagecaaagggcageccegagaaccacaggtgtacaccetgccccat 30 agagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacggctccttcttcctctacaccaacct caccgtggacaagagcaggtggcagcagggggaacgtettctcatgctccgtgatgcatgaggctctgcacaaccactacacgca gaagagcctctccctgtctccgggtaaatgatctaga

35 Amino acid sequence:

DOEPKSSDKTHTSPPSSAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHED PEVKFNWYVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCLV SNKALPAPIEKTISKAKGQPREPQVYTLPPSRDELTKNOVSLTCLVKGFYPSDIAVE WESNGQPENNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHN HYTQKSLSLSPGK

#### 22 2H7 scFv VHL11S hIgG1 (SSS-S)H K322SCH2 WCH3

Nucleotide sequence:

aagettgeegee at ggattite aagtge agattite agetteet getaate agtgette agte at aat tgee agaggae aa at tgttetet to the state of the state45 cccag totccag caat cct gtct gcatctccag gg gag aag gtcacaat gactt gcag gg ccag ctcaa gt gtaa gttacat gcact gtaccagcagaagccaggatcctcccccaaaccetggattatgccccatccaacetggettetggagtccctgctcgcttcagtg taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggagctctcaggcttatctacagcagtctggggctgagtcggtgaggcctgggggcctcagtgaagatgtcctgcaaggcttctgcc tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat 

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cagocijacatotigaagactigeggidatiticigigeaagagigigiacitatagiaactitactigatectigatgidiggigidactigacijacitacigatectigategigidacagigidacaegidacaegidaca

Amino acid sequence:

MDFQVQIPS†LISASVILARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
QQKPGSSPKPWIYAPSNLASGVPARFSGGSGTSYSLTISKVEAEDAATYYCQQWS
FNPPTFGAGTKLELKDGGGSGGGSGGGGSGGAYLQQSGAESVRPAGSK
ASGYTFTSYNMHWVKQTPRQCLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVYYSNSVWYFDVWGTGTTVTVSSODJEPKSSDK
THTSPPSSAPELLGGPSVFLFPPKPKDILMISRTPEVTCVVVDVSHEDPEVKFNWYV
DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCSVSNKALPAPIE
KTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPEN
NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS
FGK

# 23. 2H7 scFv VHL11S hIgG1 (SSS-S)H K322LCH2 WCH3

Nucleotide sequence:

nagettgeegecatggatttleaagtgeagatttteagetteetgetaateagtgetteagteataattgeeagaggaeaaattgtjetet cecagtetecageaateetgtetgeatetecaggggagaaggteacaatgacttgeagggeeageteaagtgtaagttacatgeact gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggageteteaggettatetacageagtetggggetgagteggtgaggeetggggeeteagtgaagatgteetgeaaggettetgge tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat cagcctgacatctgaagactctgeggtetatttctgtgeaagagtggtgtactatagtaactcttactggtacttegatgtetggggcac agggaccacggtcaccgtctcttctgatcaggagcccaaatcttctgacaaaactcacacatccccaccgtcctcagcacctgaact cctggggggaccgtcagtcttcctcttccccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcacatgcgtggtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagctacaagtgcctggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaagccaaagggcagcccgagaacca gegacategeegtggagtgggagagcaatgggeageeggagaacaactacaagaccacgeetecegtgetggacteegaegg ctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaatgatctaga

Amino acid sequence:

MDFQVQIFSFLLISASVIIARGQIVI SQSPAILSASPGFKVTIMTCRASSSVSYMHWY QQKPGSSPKPWIYAPSDNLASGVPARRSGGSGSTSYSLTISRVYAEDAATYYCQQWS FNPPTFGAGTKLELKDGGGSGGGGGGGGSQQAYLQQSGAESVRPGASVKMSCK ASGYTFTSYNMHWYKQTPRQGLEWIGAIYPGNGDTSYNQKIFKGKATLTVDKSSS TAYMOLSSLTSEDSAVYPCARVYYSNSYWYFDVWGTGTTYTYSSDQBFKSSDK

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THTSPPSSAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCLVSNKALPAPIE KTISKAKGQPREPQVYTLPPSRDELTKNQVSLTČLVKGFYPSDIAVEWESNGOPEN NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS PGK

#### 2H7 scFv VHL11S hIgG1 (CSS-S)H K322SCH2 WCH3 24.

Nucleotide sequence:

aagettgeegeeatggatttteaagtgeagatttteagetteetgetaateagtgetteagteataattgeeagaggaeaaattgttetet cccag to tecag caate ct g to tge at ctccag g g g agaag g teaca at g act tge ag g g ceag ctca ag t g taa g taa cat g cae t gggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggettctggagtccctgctcgcttcagtg g cagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttttaacccacccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggagctetcaggcttatctacagcagtctggggctgagtcggtgaggcctggggcctcagtgaagatgtcctgcaaggcttetggc taca catttac cagttaca at at geact gggtaa ag cagaca cct ag a cag gg cct gga at gga tt gga gct at tt at ccag ga a at ga acct ag accept ga at gga at gggtgatacticctacaatcagaagttcaagggcaaggccacactgactgtagacaaatcctccagcacagcctacatgcagctcag cagectgacatctgaagactctgcggtctatttctgtgcaagagtggtgtactatagtaactcttactggtacttcgatgtctggggcac agggaccacggtcaccgtctcttctgatcaggagcccaaatcttgtgacaaaactcacacatccccaccgtcctcagcacctgaact cetggggggacegteagtetteetetteeceecaaaacecaaggacacecteatgateteeeggacecetgaggteacatgegtgg tggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagc cgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggag tacaagtgctcggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaagccaaagggcagccccgagaacca gcgacatcgccgtggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacgg ctecttetteetetacageaageteacegtggacaagageaggtggeageaggggaaegtetteteatgeteegtgatgcatgagg etetgeacaaccactacacgcagaagagcetetecetgteteegggtaaatgatetaga

Amino acid sequence:

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY OOKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS FNPPTFGAGTKLELKDGGGSGGGGGGGGGSSQAYLQQSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS TAYMOLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSCDK THTSPPSSAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCSVSNKALPAPIE 35 KTISKAKGOPREPOVYTLPPSRDELTKNOVSLTČLVKGFYPSDIAVEWESNGOPEN NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS PGK.

#### 40 25. P331S CH2

Nucleotide sequence:

cetgaacteetggggggacegteagtetteetetteeceecaaaaceeaaggacaceeteatgateteeeggaceeetgaggteae gacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatg gcaaggagtacaagtgcaaggtetecaacaaageceteccagectecategagaaaacaatetecaaagecaaa

Amino acid sequence PELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEVHN AKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPASIEKTISKAK

26. P331S CH2 WCH3

Nucleotide sequence:

10 Amino acid sequence

15

PELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEVHN AKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPASEKTISKAKG REPEQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPPV LDSDGSFFLYSKLTVDKSRWOOGNVFSCSVMHEALHNHYTOKSLSLSPGK

EDDOD'T ETSKET VDKSKW QQOTVYSCS VMILEALITHTT I QKSLSESTOK

# 2H7 scFv VH L11S (SSS-S)H P331S CH2 WCH3

Nucleotide sequence:

aagettgeegee atggatttte aagtge agatttte agetteet get aat eagtgette agte at aattgee agaggae aa attgtteet the aget agaggae agatt to the state of the20 cccagtctccagcaatcctgtctgcatctccaggggagaaggtcacaatgacttgcagggccagctcaagtgtaagttacatgcact gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggageteteaggettatetacageagtetggggetgagteggtgaggeetggggeeteagtgaagatgteetgeaaggettetgge 25 tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat gglgatacttectaeaateagaagtteaagggeaaggeeacaetgaetgtagacaaatecteeageacageetacatgeageteag cagcctgacatctgaagactctgcggtctatttctgtgcaagagtggtgtactatagtaactcttactggtacttcgatgtctggggcac agggaccacggtcaccgtctcttctgatcaggagcccaaatcttctgacaaaactcacacatccccaccgtcctcagcacctgaact cctggggggaccgtcagtcttcctcttcccccaaaacccaaggacacctcatgatctcccggacccctgaggtcacatgcgtgg 30 tggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagc cgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggag tacaagtgcaaggtctccaacaaagccctcccagcctccatcgagaaaacaatctccaaagccaaagggcagccccgagaacca caggtgtacaccetgeccecatecegggatgagetgaccaagaaccaggteagcetgacetgectggteaaaggettetatecea gcgacatcgccgtggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacgg 35 ctecticttcctctacagcaagctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgagg ctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaatgatctaga

Amino acid sequence

MDFQVQIFSFLLISASVILARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
40 QQKPGSSPKPWIYAPSNLASGVPARPSGGSGTSYSLTISRVEAEDAATYYCQQWS
FNPPTFGAGTKLELKDGGGSGGGSGGGGSQQAYLQQSGAESVRPGASVKMSCK
ASGYTFTSYNMHWYKQTPRQGLEWIGAIYPGNGDTSYNQKEKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSSDK
THTSPPSSAPELLGGPSVFLPPKPKDTLMISRTPEVTCVVDVSHEDPEVKFNWYV

45 DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPASIE KTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPEN NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS PGK

Nucleotide sequence:

aagettgeegeeatggatttteaagtgeagatttteagetteetgetaateagtgetteagteataattgeeagaggaeaaattgttetet cccagtctccagcaatcctgtctgcatctccaggggagaaggtcacaatgacttgcagggccagctcaagtgtaagttacatgcact ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggttggatctggaggaggtg ggageteteaggettatelacageagtetggggetgagteggtgaggeetggggeeteagtgaagatgteetgeaaggettetgge tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat 10 cagoctgacatctgaagactctgcggtctatttctgtgcaagagtggtgtactatagtaactcttactggtacttcgatgtctggggcac agggaccacggtcaccgtctcttctgatcaggagcccaaatcttgtgacaaaactcacacatccccaccgtcctcagcacctgaact cctggggggaccgtcagtcttcctcttccccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcacatgcgtgg cgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggaq 15 tacaagtgcaaggtctccaacaaagccctcccagcctccatcgagaaaacaatctccaaagccaaagggcagccccgagaacca caggtgtacaccetgeccccatccegggatgagetgaccaagaaccaggteagcetgacetgectggtcaaaggettetatecca gcgacatcgccgtggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacgg ctcettetteetetacageaageteacegtggacaagageaggtggeageaggggaacgtetteteatgeteegtgatgeatgagg ctetgeacaaccactacacgeagaagagceteteeetgteteegggtaaatgatetaga

20 Amino acid sequence

MDFQVQJFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY QQKPGSSPKPWTAPSNLASGVPARFSGSGGTSYSLTISRVEAEDAATTYYCQQWF FNPPTFGAGTKLEILKDGGGSGGGGSGGGGSSQAYLQOSGAESVRPGASVKMSCK

- 25 ASGYTFTSYNMHWVKQTPRQGLEWIGALTPGNGDTSYNQKFKGKATLTVDKSSS
  TAYMQLSSLTSEDSAVYFCARVVYYSNSVWYFDVWGTGTTVTVSSDQEPKSČDK
  THTSPPSSAPELLGGPSVFLFPFKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV
  DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPASIE
  KTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPEN
  NVKTTPPUVLDSRGSEFI VSKLTVDKSBWGGSNVFSCSVAME AL HANHETOR EJ SL
- 30 NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS PGK

# 29. T256N CH2 region

- Nucleotide sequence:
- 40 Amino acid sequence

PELLGGPSVFLFPPKPKDTLMISRNPEVTCVVVDVSHEDPEVKFNWYVDGVEVHN AKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAK

# T256N CH2 WCH3

45 Nucleotide sequence:

50

actocgacggetecttettectetaeageaageteaeegtggacaagaggaggtggeaggggaaegtetteteatgeteegtg atgeatgaggetetgeaeaaceaetaeaegeagaagageeteteeetgteteegggtaaatgatetaga

Amino acid sequence

5 PELLGGPS VFLFPPKPKDTLMISRNPEVTCVVVDVSHEDPEVKFNWYVDGVEVHN AKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALP APIEKTISKAKG QPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVE WESNGQPENNYKTTPPV LDSDGSFFLYSKLTVDKSRWQOGNVFSCSVMHEALHNHYTOKSLSLSPGK

# 10 31. 2H7 scFv VH L11S (SSS-S)H T256N CH2 WCH3

Nucleotide sequence:

aagettgeegeeatggatttteaagtgeagatttteagetteetgetaateagtgetteagteataattgeeagaggaeaaattgttetet cccagtctccagcaatcctgtctgcatctccaggggagaaggtcacaatgacttgcagggccagctcaagtgtaagttacatgcact 15 ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat20 ggtgatacttectacaateagaagtteaagggeaaggeeacaetgaetgtagacaaateeteeageacageetacatgeageteag cagcctgacatctgaagactctgcggtctatttctgtgcaagagtggtgtactatagtaactcttactggtacttcgatgtctggggcac agggaccacggtcaccgtctcttctgatcaggagcccanatcttctgacanaactcacacatccccaccgtcctcagcacctgaact cctggggggaccgtcagtcttcctcttcccccaaaacccaaggacaccctcatgatctcccggaaccctgaggtcacatgcgtgg tggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagc 25 egegggaggagcagtacaacagcaegtacegtgtggtcagegtectcacegtectgcaecaggaetggetgaatggcaaggag tacaagtgcaaggtetecaacaaageceteccagececcategagaaaacaatetecaaagecaaagggcageceegagaace acaggtgtacaccetgccccatcccgggatgagctgaccaagaaccaggtcagcetgacetgcetggtcaaaggettetatccc agcgacategeegtggagtgggagagcaatgggeageeggagaacaactacaagaccaegeeteeegtgetggacteegaeg geteettetteetetacageaageteacegtggacaagageaggtggcagcaggggaacgtettetcatgeteegtgatgcatgag 30 getetgeacaaceactacaegeagaagageeteteeetgteteegggtaaatgatetaga

Amino acid sequence

MDFQVQIPSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS
SPRPPTFGAGTKLELIKDGGGSGGGGGGGGSGGASQAYLQQSGAESVRPGASVKMSCK
ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYTCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSSDK
THTSPPSSAPELLGGPSVTLPPPKDTLMISRNPEVTCVVVDVSHEDPEVKFNWY
VDGVEVIINAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPI
EKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPE
NNYKTTPPVLDSDGSFFLYSKLTVDKSRWOOGNVPSCSYMHEALHNHYTOKSLSL

# 32. 2H7 scFv VH L11S (CSS-S)H T256N CH2 WCH3

Nucleotide sequence:

SPGK

45

50

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ggagctetcaggettatetaeageagtetggggetgagteggtgaggeetggggeeteagtgaagatgteetgeaaggettetgge tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat ggtgataetteetacaateagaagtteaagggeaaggeeacaetgactgtagacaaateeteeageacageetacatgeageteag cagcet gacatet gaaga actet geggtet att tet gt geaagagt ggt gt actatagt aactet tact ggt act te gat gt et ggg geaecctggggggaccgtcagtcttcctcttccccccaaaacccaaggacaccctcatgatctcccggaaccctgaggtcacatgcgtgg tggtggacgtgagocacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagc tacaagtgcaaggtetecaacaaageceteecageececategagaaaacaatetecaaagecaaagggcageecegagaaec agegaeategeegtggagtgggagageaatgggeageeggagaacaactacaagaccaegeeteeegtgetggaeteegaeg getetgeacaaccactacaegcagaagageeteteeetgteteegggtaaatgatetaga

#### 15 Amino acid sequence

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQQWS FNPPTFGAGTKLELKDGGGSGGGGSGGGSSQAYLQQSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS TAYMOLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSCDK

20 THTSPPSSAPELLGGPSVFLFPPKPKDTLMISRNPEVTCVVVDVSHEDPEVKFNWY VDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPI EKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPE NNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSL 25 SPGK:

#### 33. RTPE/QNAK (255-258) CH2

Nucleotide sequence:

cctgaactcctggggggaccgtcagtcttcctcttccccccaaaacccaaggacaccctcatgatctcccagaacgctaaggtcac 30 gacaaagccgcgggaggaggagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggeaaggagtacaagtgeaaggteteeaaeaaageeeteecageeeecategagaaaaaaateteeaaageeaaa

Amino acid sequence

PELLGGPSVFLFPPKPKDTLMISONAKVTCVVVDVSHEDPEVKFNWYVDGVEVHN 35 AKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAK

# RTPE/QNAK (255-258)CH2 WCH3

Nucleotide sequence:

40 gacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggagtacaagtgcaaggtetccaacaaagceetceeagceeceategagaaaacaatetccaaagceaaagggcagece cttctatcccage gacatege egt ggag t gg gag ag ea at gg gcage eg gag aa caacta caa gac cae geet cee gt get ggag ag accae gag accae geet consistent of the consistency 5

actecgacggeteettetteetetaeageaageteacegtggacaagageaggtggeageagggaaegtetteteatgeteegtg atgeatgaggetetgeacaaccactacaegeagaagageeteteeetgteteegggtaaatgatetaga

Amino acid sequence

PELLGGPSVFLFPPKPKDTLMISQNAKVTCVVVDVSHEDPEVKFNWYVDGVEVHN 50 AKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAKG

10

15

20

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 $\label{thmodel} QPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPPV\\ LDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSPGK$ 

# 35. 2H7 scFv VH L11S (SSS-S)H RTPE/QNAK (255-258)CH2 WCH3

Nucleotide sequence:

aagettgeegecatggatttteaagtgeagatttteagetteetgetaateagtgetteagteataattgeeagaggaeaaattgttetet cccagtctccagcaatcctgtctgcatctccaggggagaaggtcacaatgacttgcagggccagctcaagtgtaagttacatgcact ggtaccagcagaagccaggatcctcccccaaaccetggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggageteteaggettatetaeageagtetggggetgagteggtgaggeetggggeeteagtgaagatgteetgeaaggettetgge tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat ggtgatacttectacaateagaagtteaagggeaaggeeacaetgaetgtagacaaateeteeageacageetacatgeageteag cagcctgacatctgaagactctgcggtctatttctgtgcaagagtggtgtactatagtaactcttactggtacttcgatgtctggggcac agggaccacggtcaccgtctcttctgatcaggagcccaaatcttctgacaaaactcacacatccccaccgtcctcagcacctgaact cctggggggaccgtcagtcttcctcttccccccaaaacccaaggacaccctcatgatctcccagaacgctaaggtcacatgcgtgg tggtggacgtgagccacgaagaccetgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagc cgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggag tacaagtgcaaggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaagccaaagggcagcccgagaacc agegacategeegtggagtgggagageaatgggeageeggagaacaactacaagaeeaegeeteeegtgetggacteegaeg geteettetteetetacageaageteacegtggacaagageaggtggcageagggggaacgtetteteatgeteegtgatgeatgag getetgeacaaceactacaegeagaagageeteteeetgteteegggtaaatgatetaga

25

Amino acid sequence

MDFQVQIESFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY
QVKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISKVEAEDAATYYCQQWS
FNPPTFGAGTKLELKDGGGSGGGGSGGGGSSQAYLQQSGAESVRPGASVKMSCK

ASGYTFTSYNMHWVKQTPRQCLEWIGATPPGNGDTSYNQKFKGKATLTVDKSSS
TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSSDK
THTSPPSSAPELLGGPSVBLPPKPKDTLMISQNAKVTCVVVDVSHEDPEVKFNWY
VDGVEVHNAKTKPREGYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKATPAPI
EKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPE
NNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSL
SPGK

# 36. 2H7 scFv VH L11S (CSS-S)H RTPE/QNAK (255-258)CH2 WCH3 Nucleotide sequence:

cgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggag tacaagtgcaaggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaagccaaagggcagcccgagaacc acagglgtacaccetgececcatecegggatgagetgaccaagaaccaggtcagectgacetgeetggtcaaaggettetatece agcgacategccglggagtgggagagcaatgggeagccggagaacaactacaagaccacgcetecegtgetggactecgacg gctccttcttcctctacagcaagctcaccgtggacaagagcaggtggcagcagggggaacgtcttctcatgctccgtgatgcatgag getetgeacaaccactacaegeagaagageeteteeetgteteegggtaaatgatetaga

#### Amino acid sequence

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY OOKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCOOWS FNPPTFGAGTKLELKDGGGSGGGGGGGGGSOAYLOOSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKOTPROGLEWIGAIYPGNGDTSYNOKFKGKATI TVDKSSS TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDOEPKSCDK THTSPPSSAPELLGGPSVFLFPPKPKDTLMISONAKVTCVVVDVSHEDPEVKFNWY 15 VDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPI EKTISKAKGOPREPOVYTLPPSRDELTKNOVSLTCLVKGFYPSDIAVEWESNGOPE NNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSL

#### 20 36. K290Q CH2 region

SPGK

25

30

### Nucleotide sequence:

ectgaactcetggggggacegtcagtcttcctcttcccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcac gacacageegegggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatg gcaaggagtacaagtgcaaggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaagccaaa

Amino acid sequence:

PELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEVHN AKTQPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAK

#### 37. K290Q CH2 WCH3

### Nucleotide sequence:

Cetgaacteetggggggacegteagtetteetetteeecceaaaacceaaggacacceteatgateteeeggaceeetgaggtea catgogtggtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgcca 35 agacacagcegegggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtctgcaccaggactggetgaat ggcaaggagtacaagtgcaaggtetecaacaaageeeteecageeeecategagaaaacaatetecaaagecaaagggcagee gcttctateccagegacategecgtggagtgggagagcaatgggcageeggagaacaactacaagaccaegeeteecgtgetg gactecgaeggeteettetteetetaeageaageteaeegtggaeaagageaggtggeageaggggaaegtetteteatgeteegt 40 gatgcatgaggetetgcacaaccactacacgcagaagagcetetecetgtetecgggtaaatgatetaga

### Amino acid sequence:

PELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEVHN AKTOPREEOYNSTYRVVSVLTVLHODWLNGKEYKCKVSNKALPAPIEKTISKAKG QPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTPPV

45 LDSDGSFFLYSKLTVDKSRWOOGNVFSCSVMHEALHNHYTOKSLSLSPGK

# 2H7 scFv VH L11S (SSS-S)H K290Q CH2 WCH3

# Nucleotide sequence:

 $50 \quad \text{aagettgeegecatggatttteaagtgeagatttteagetteetgetaateagtgetteagteataattgeeagaggacaaattgttetet}$ cccag to to cag caate ct gtot geat ctccag gg gag aag gto acaat gact t g cag gg ccag ctcaa gt gtaa gt ta cat geat ctcag gag aag a

15

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ggtaccagcagaagccaggatcctccccaaaccctggatttatgccccatccaacctggettetggagtccctgctcgetteagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgtteggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggagctctcaggcttatctacagcagtctggggctgagtcggtgaggcctggggcctcagtgaagatgtcctgcaaggcttctggc tacacatttaccagttacaatatgcactgggtaaagcagacactagacagggcctggaatggattggagctatttatccaggaaat cagcctgacatctgaagactctgcggtctatttctgtgcaagagtggtgtactatagtaactcttactggtacttcgatgtctgggcac agggaccacggtcaccgtctcttctgatcaggagcccaaatcttctgacaaaactcacacatccccaccgtcctcagcacctgaact cctgggggaccgtcagtcttcctcttcccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcacatgcgtgg tggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacacagc cgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggag tacaagtgcaaggtetecaacaaageeeteecageeeecategagaaaacaatetecaaaggcaaagggcageeeegagaace agcgacalcgccglggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacg gctccttcttcctctacagcaagctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgag getetgeacaaceactacaegeagaagageeteteeetgteteegggtaaatgatetaga

# Amino acid sequence:

- MDFOVOIFSFLLISASVIIARGOIVLSOSPAILSASPGEKVTMTCRASSSVSYMHWY 20 OOKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCOOWS FNPPTFGAGTKLELKDGGGSGGGGGGGGGSOAYLOOSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKOTPROGLEWIGAIYPGNGDTSYNOKFKGKATLTVDKSSS TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDOEPKSSDK THTSPPSSAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV DGVEVHNAKTOPREEOYNSTYRVVSVLTVLHODWLNGKEYKCKVSNKALPAPIE
- 25 KTISKAKGOPREPOVYTLPPSRDELTKNOVSLTCLVKGFYPSDIAVEWESNGOPEN NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS PGK.

#### 30 39. 2H7 sefv VH L11S (CSS-S)H K290O CH2 WCH3

### Nucleotide sequence:

aagettgeegecatggatttteaagtgeagatttteagetteetgetaateagtgetteagteataattgeeagaggaeaaattgttetet cccagtetecageaateetgtetgeatetecaggggagaaggteacaatgaettgeagggeeageteaagtgtaagttaeatgeaet ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtcctgctgcttcagtg 35 gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggagctctcaggcttatctacagcagtctggggctgagtcggtgaggcctggggcctcagtgaagatgtcctgcaaggcttctggc tacacatttaccagttacaatatgcactgggtaaagcagacactagacagggcctggaatggattggagctatttatccaggaaat 40 cagectgacatetgaagactetgeggtetatttetgtgcaagagtggtgtactatagtaactettaetggtaettegatgtetggggeae agggaccaeggteaccgtetettetgateaggageceaaatettgtgacaaaacteacacateccacegteeteageacctgaact cctggggggaccgtcagtcttcctcttcccccaaaacccaaggacacctcatgatctcccggacccctgaggtcacatgcgtgg tggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacgtggaggtgcataatgccaagacacagc cgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggag

45 tacaagtgcaaggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaaggcaaagggcagcccgagaacc agcgacatcgccgtggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacg gctccttcttcctctacagcaagctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgag getetgeacaaceactacaegeagaagageeteteeetgteteegggtaaatgatetaga

Amino acid sequence:

PCT/US2003/041600

MDFOVOIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY OOKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCOOWS FNPPTFGAGTKLELKDGGGSGGGGSGGGSSQAYLQQSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS TAYMOLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSCDK THTSPPSSAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV DGVEVHNAKTOPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIE KTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPEN NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS 10 PGK

#### 40. A339PCH2

Nucleotide sequence:

cetgaacteetggggggacegteagtetteetetteececcaaaacccaaggacacceteatgateteeeggacceetgaggteae 15 gacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatg gcaaggagtacaagtgcaaggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaacccaaa

Amino acid sequence:

PELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEVHN 20 AKTKPREEOYNSTYRVVSVLTVLHODWLNGKEYKCKVSNKALPAPIEKTISKPK

#### 41. A339P/CH2 WCH3

25 Nucleotide sequence:

cctgaactcetgggggaccgtcagtcttcctcttcccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcac, gacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatg gcaaggagtacaagtgcaaggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaacccaaagggcagccc cttctatcccagcgacatcgccgtggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggatgcatgaggctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaatgatctaga

35 Amino acid sequence:

40

PELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEVHN AKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKPKG OPREPOVYTLPPSRDELTKNOVSLTCLVKGFYPSDIAVEWESNGOPENNYKTTPPV LDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSPGK

#### 2H7 scFv VHL11S (SSS-S)H A339P CH2 WCH3 42.

Nucleotide sequence:

a agett geograat gg at ttt caagt ge ag at ttt cagett cet get aat cagt get te ag te at aat t ge cag ag ga caa at t g tt e te te te geografie to get general to get geografie to get geografie to get general to general to get general to general to get general to general to general to get general to get general to general to get general to general to get general to gcccagtetecageaatectgtetgeatetecaggggagaaggteacaatgaettgeagggeeageteaagtgtaagttaeatgeaet ggtaccag cag gate ctcccccaaaccet ggatttat gecccate caacct ggettet ggag tee et gete ag type of the contract of the contract grant g45 g cagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgcaccttattactgccagcagtggagtttactgccacttattactgccagcagtggagtttactgcacttattactgccagcagtggagtttactgcacttattactgccagcagtggagtttactggaggctgaagatgctgcaccttattactgccagcagtggagtttactgcacttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggtttactgcacttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgcaccttattactgccagcagtggaggttgaagatgctgaagatgaagatgctgaagata acc caccae get c g g t g c t g g accaa g c t g a g c t g a a g a t g g c t g g c t g g t g g t c t g g a g g a g g t g a t c t g g a g g a g g t g g a t c t g g a g g a g g t g g a t c t g g a g g a g g t g g a t c t g g a g g a g g t g g a t c t g g a g g a g g t g g a t c t g g a g g a g g t g g a t c t g g a g g a g g t g g a c c a a g a t c t g a g a g a g g a g g t g g a c c a a g a c a a c a a a c a a c a a c a a c a a c a a c a a c a a c a a c a a c a a c a a c a a c a a c a a c a ataca catttac cagttaca at at geactgggta a ag cag acacctaga cag ggcct gga at gga at t gga gct at ttat ceag ga a at the cagtaga cagtaga at the gtgata et tecta caat caga ag tt caa ggge aa gge caca ct gact gtaga caaat cet ceage acage ctae at geage teager ag the comparison of the50 

### PCT/US2003/041600

agggaccacggtcaccgtctcttctgatcaggagcccaaatcttctgacaaaactcacacatccccaccgtcctcagcacctgaact cetggggggaccgtcagtcttcctcttcccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcacatgcgtggta ca agt g ca ag g to teca a ca a ag e cote coag e coccate g aga a a accate teca a accea a ag g g ca g cocc g ag a accea and g constraints and the constraints are the constraints are the constraints and the constraints are gegacategeegtggagtgggagageaatgggeageeggagaacaactacaagaccaegeeteeegtgetggacteegaeg get cette tte et ctacage aa get caceg t g gacaag aa get g get ag ea geg gaa e get et te te at get ee get gat geat gaa gegen gegen gedeuwerd.gctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaatgatctaga

10

20

Amino acid sequence: MDFOVOIFSFLLISASVIIARGOIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY QOKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQOWS FNPPTFGAGTKLELKDGGGSGGGSGGGSSQAYLQQSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKQTPROGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS 15 TAYMOLSSLTSEDSAVÝFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSSDK THTSPPSSAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIE KTISKPKGQPREPQVYTLPPSRDELTKNQVSLTČLVKGFYPSDIAVEWESNGQPEN NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS PGK

#### 2H7 scFv VHL11S (CSS-S)H A339P CH2 WCH3 43.

25 Nucleotide sequence: a agett geogecatgg at tit caagt geag at tit cagetteet get a at cagt get teag reat a at tig ceag agg acaa at tig the total content of the cocccag to to cag caat cot g to to cag g g g a g a g g to a caat g a ctt g cag g g ccag ct caa g t g taa g t a cat g cact g cag g cag ct caa g t g cact g cact g cac g ct caa g t g cact g cac g ct caa g t g cac g ct cac gggtaccag cag a age caggate ctcccccaa accet ggatt tatgccccatccaacct ggettet ggag tecet gete gette age ggattet ggag tecet get get ggattet ggag tecet get ggattet ggag tecet ggagg cag t g g g acctett actete teaca at cag cag a g t g g a g et g a g at g et g e caet t at tact g e cag t g g a g t t t a caet a g e caet a gggagctctcaggcttatctacagcagtctggggctgagtcggtgaggcctgggggcctcagtgaagatgtcctgcaaggcttctggc tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaat cagcet gacatet gaagactet geggtet att tet gtg caagagt gg tg tactatagt aactet tact gg tact te gat gt et gg gg cacatet gaagactet gaagactet gat gegeg cacatet gaagactet gggaccacggtcaccgtctcttctgatcaggagcccaaatcttgtgacaaaactcacacatccccaccgtcctcagcacctgaact 35 cctggggggaccgtcagtcttcctcttccccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcacatgcgtggta can agt g caa ag g to to caa caa ag g coccea ag e cocca to g ag a aa acaa to to caa accea aag g g cag cocc g ag a accea ag g cag cocca ag ag accea ag g cag cocca ag ag accea ag g cag cocca ag ag accea ag accea ag ag accea ag accea ag accea ag accea ag ag accea ag accea ag ag accea ag acces a40 agcgacatcgccgtggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacg geteettetteetetacageaageteacegtggacaagageaggtggcagcaggggaaegtetteteatgeteegtgatgeatgag gctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaatgatctaga

45 Amino acid sequence:

MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQOWS FNPPTFGAGTKLELKDGGGSGGGGGGGGGGSOAYLQQSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSCDK 50 THTSPPSSAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV

PCT/US2003/041600

DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIE KTISKPKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPEN NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS PGK

5

# 44. G28-1VH

Nucleotide sequence:

gegticeagetgenageagtitgsactigsactigsactigsgetteagtgaagatticetgenaggettetgttactatit 10 actggctacantatganetgggtgaageagantaatggaangaagecttgagtgattgsanatatatgstetgatat cotacaaceggnagticaagggcaaggccaattgactgtagacaatcotccagcacagoctacatgcagctcangagtetgac atctgaggactctgcagtctattactgtgcaagatoggteggccctatggactactggggtcaaggaacetcagtcacegtctcttct gatcag

15 Amino acid sequence:

AVQLQQSGPĒLEKPGASVKISCKASGYSFTGYNMNWVKQNNGKSLEWIGNIDPY YGGTTYNRKFKGKATLTVDKSSSTAYMQLKSLTSEDSAVYYCARSVGPMDYWG OGTSYTVSSDO

### 20 45, G28-1VL

Nucleotide sequence:

Amino acid sequence:

30 MVSTAQFI.GLLLLWLTGGRCDIQMTQSPASLSASVGETVTITCRTSENVYSYLAW YQQKQGKSQLUSFAKTLAEGYPSRFSGSGSGTQFSLKISSLQPEDSGSYFCQHHS DNPWTGGGTELEIKGGGSGGGGSGGGGSS

### 35 46. G28-1 scFv

Nucleotide sequence:

ttctgatcag

Amino acid scquence:

MVSTAQFLGILLLWLTGGRCDIQMTQSPASLSASVGETVTITCRTSENVYSYLAW 70QKQGKSPQLLVSFAKTLAEGVPSRFSGSGSGTQFSLKISSLQPEDSGSYFCQHHS DNPWTFGGGTELEIKGGGSGGGGSGGGGSSAVQLQQSGPELEKPGASVKISCKA

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 $SGYSFTGYNMNWVKQNNGKSLEWIGNIDPYYGGTTYNRKFKGKATLTVDKSSST\\ AYMQLKSLTSEDSAVYYCARSVGPMDYWGQGTSVTVSSDQ\\$ 

# 5 47. G28-1 VHL11S

Nucleotide sequence:

geggtocagedgeageagtitggacetgagetggaaagectgggeggtotggtgagtgagtatteetgeaaggetttegtttacteatte actggetacaatatgaactgggtgaageagaataatggaaagageettgagtggattggaaatattgateettattatggtgatata cetacaaceggaagtteaagggeaaggecaeattgactgtagacaaatoctecageacageetacagcagetcaaggtetgae atetgaggacetgeagtetattactgtgeagateggeegeeetatggactactggggteaaggaacetcagtcacegtetetet atetgaggacetgeagtetattactgtgeagateggeegeeetatggactactggggteaaggaacetcagtcacegtetettet ateca

### Amino acid sequence:

15 AVQLQOSGPESEKPGASVKISCKASGYSFTGYNMNWVKQNNGKSLEWIGNIDPYY GGTTYNRKFKGKATLTVDKSSSTAYMQLKSLTSEDSAVYYCARSVGPMDYWGQ GTSVTVSSDQ

# 20 48. G28-1 VHL11S scFv

Nucleotide sequence;

# Amino acid sequence:

ttctgatcag

MVSTAQFLĞILLLWLTGGRCDIQMTOSPASILSASVGETVTITCRTSENVYSYLAW
35 YQQKQĞKSPQLLVSFAKTLAEGVPSRFSGSGSGTQFSLKISSLQPEDSGSYFCQHHS
DNPWTFGGGTELEIKGGGSGGGGSGGGGSSAVQLQQSGPESEKPGASVKLISCKA
SGYSFTGYNMNWVKQNNGKSLEWIGNIDPYYGGTTYNRKFKGKATLTVDKSSST
AYMQLKSLTSEDSAVYYCARSVCPMDYWGGGTSVTVSSDO

# 40 49. G28-1 scFv (SSS-S)H WCH2 WCH3

Nucleotide sequence:

PCT/US2003/041600

WO 2005/017148

agtottectottecececanaacecaaggacacectatgateleceggacecetgaggicacatgottgstgagetgagetgagec acgangacectgaggicangtranatggtacgtgaeggergegaggtgetaatgecaagacaagacaagecggggaggagea gtacaacageacgtacegtgtgetaggctectacecgtotgacecaggactggetgaatggcaaggaagacaggtgcaaggt tecaacaaagecetecagececategagaaanacaatetecanagecaagggeageceggaaaceacaggtgtacacect gececentecegggatgaegtgaccaagaacagatgagetgacetgtgctgctggtgaaaggettatetaceagggatacacet gagtggggagetaatgggcageaggagaaacactacaagacacagectocgtggtagaaggettatetaceaggagatacagect agcaagetaacegtggacaagaagaggggaaggagggggagttetteatggtetggtagacgatggagettettetteetaa gacaagetaacegtggacaagaagaggggaagaagatataga

### 10 Amino acid sequence:

15

20

MVSTAQFLGLLLLWLTGGRCDIQMTQSPASLSASVGETVTITCRTSENVYSYLAW
YQQKQGKSPQLLVSFAKTLAEGYPSKFSGSGSGTQFSLKISSLQPEDSGSYFCQHHS
DNFWTFGGGTBLEEK GGGSGSGGGSGGGGSSAVQLQQSGFELEKPGASVKISCKA
SGYSFTGYNMNWVKQNNGKSLEWIGNIDPYYGGTTYNRKFKGKATLTVDKSSST
AYMQLKSLTSEDSAVYYCARSVGFMDYWGQGTSVTVSSDHDQEPKSSDKTHTSP
PSSAPELLGGFSVFLFPFPKBVDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVE
VHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISK
AKGQPREPQVTLPPSRDELTKNQVSLTCLVKGFYPSDLAVEWESNGQPENNYKT
TPPYLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSFGK

50. G28-1 scFv IgAW H IgG1WCH2 WCH3

Nucleotide sequence: a agett geograat gglate cacaget cagt teett gggt t get get ggctac aggt ggcag at gt gacate cag at gacter and the second sagtote cage ctccct at ctg catctg tgg gag agactg tcaccat cacat gtc gaa caa gtgaa aatgtt tacagtt at ttgg ctt ggt25 gtggatcaggcacacagttttctctgaagatcagcagcctgcagcctgaagattctggaagttatttctgtcaacatcattccgataat cg t cag cgg to cag ctg cag cag tctg gacctg ag ctg gaaa ag cct gg cgctt cag tgaag at tt cct gcaag gctt ctgg tt act gacctg gaaa ag cct gg gaaa ag ctt gag gacctg gaaa ag cct gg gaaa ag ctt gag gaaa ag cct gg gaaa ag cct gag gaaa ag cct gacatteactggctacaatatgaactgggtgaagcagaataatggaaagagccttgagtggattggaaatattgatccttattatggtggtgaatggaaatattgatccttattatggtggtgaatggaaatattgatccttattatggtggtgaatggaaatattgatccttattatggtggtgaatggaactacctacaaccggaagttcaagggcaaggccacattgactgtagacaaatcctccagcacagcctacatgcagctcaagagtct 30 gacatet gaggactet geagtet attact gtgeaa gateg gteggeect at ggactact ggggteaa ggaacctea gteacc gtete gaggactet gaggttetgateageeagtteeeteaacteeacetaeceeateteecteaacteeacetaeceeateteecteatgegeacetgaacteetgg ggggaccgtcagtcttcctcttccccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcacatgcgtggtggtg gacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagccgcg ggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggagtacaa 35 gtgcaaggtetecaacaaageeeteecageeeceategagaaaacaatetecaaageeaaagggcageeeegagaaceacagg categeegtggagtgggagagcaatgggcagceggagaacaactacaagaccaegceteeegtgetggacteegaeggeteet tottoctotacagcaagctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaatgatctaga 40

Amino acid sequence:

MVSTAQFLGLLLWLTGGRCDIQMTQSPASI.SASVGETVTITCRTSENVYSYLAW
YQQKQGKSPQLLVSFAKTI.AEGVPSRFSGSGSTQFSLKISSLQPEDSGSYFCQHHS
45 DNPWTFGGGTELEIKGGGGSGGGGGGGSGAVQLQQSGPELEKPGASVKISCKA
SGYSFTGYNMNWVKQNNGKSLEWIGNIDPYYGGTTYNRKFKGKATLTVDKSSST
AYMQLKSLTSEDSAVYYCARSVOPMDV WQGTSVTVSDQPVPSTPPFPSPTPPF
PSPSCAPELLGGPSVFLFPKPKDTLMISRTPEVTCVVDVSHEDPEVKFNWYVDG
VEVHNAKTKPRIEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTI
50 SKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNY

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KTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSPG K

# 51. G28-1 scFv VHL11S (SSS-S)H WCH2 WCH3

5 Nucleotide sequence:

aagettgeegecatggtatecacageteagiteettgggttgetgetgetgtggettacaggtggeagatgtgacatecagatgacte agtetecageetecetatetgeatetgtgggagagaetgteaceateacatgtegaacaagtgaaaatgtttacagttatttggettggt atcagcagaaacagggaaaatctcctcagctcctggtctcttttgcaaaaaccttagcagaaggtgtgccatcaaggttcagtggca gtggatcaggcacacagttttctctgaagatcagcagcctgcagcctgaagattctggaagttatttctglcaacatcattccgataatcgtcagcggtccagctgcagcagtctggacctgagtcggaaaagcctggcgcttcagtgaagatttcctgcaaggcttctggttact catteactggctacaatatgaactgggtgaagcagaataatggaaagagcettgagtggattggaaatattgateettattatggtggt actacctacaaceggaagttcaagggcaaggccacattgactgtagacaaatcctccagcacagcctacatgcagctcaagagtct gacatetgaggactetgeagtetattaetgtgeaagateggteggeectatggactaetggggteaaggaaceteagteaeegtete ttetgateaggageceaaatettetgacaaaacteacacateeceacegteeteageacetgaacteetggggggacegteagtett cetettececeaaaacccaaggacacceteatgateteceggacccetgaggtcacatgegtggtggtggacgtgagccacgaa gaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagccgcgggggggaggagtacaa aaageceteecageececategagaaaacaatetecaaagecaaagggcageceegagaaccacaggtgtacaccetgeecee gagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacggctccttcttcctctacagcaag ctcaccgtggacaagagcaggtggcagcaggggaacgtetteteatgeteegtgatgcatgaggetetgeacaaccactacaege agaagageeteteeetgteteegggtaaatgatetaga

25 Amino acid sequence:

MVSTAQELGILLI.WI.TGGRCDIQMTQSPASISASVGETVTITCRTSENVYSYLAW
YQQKQKSPQLLVSFAKTLAEGVYSRFSGSGSGTQFSLKISSLQEDSGSYFCQHHS
DNPWTFGGGTELEIK.GGGSGGGGSGGGGSSAVQLQQSGPESEKPGASVKISCKA.
SGYSFTGYYMNWVKQNNGKSLEWIGNIDPYYGGTTYNRKFKGKATLTVDKSSST

30 AYMQLKSLTSEDSAVYYCARSVGMDYWQGGTSVYVSSDHDQEPKSSDKTHTSP
PSSAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVE
VHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPEKTISK
AKQPPEPQVYTLPPSRDELTKNQVSLTCL.VKGFYPSDJAVEWESNGGPENNYKT
TPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLSPGK

52. G28-1 scFv VHL11S (CSS-S)H WCH2 WCH3

Nucleotide sequence: a agent per la contractiva de  contractiva del contractiva de la contractiva de l

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Amino acid sequence:

MVSTAQFLGLLLWLTGGRCDIQMTQSPASLSASVGETVTITCRTSENVYSYLAW YQQKQGKSPQLLVSFAKTLAEGYPSRISGSGSGTQFSLKISSLQPEDSGSYFCQHHS DNPWTFGGGTELLER GGGGSGGGGGGGGGSAVQLQQSGFPSEKPGASVKISCKA SGYSFTGYNMNWVKQNNGKSLEWIGNIDPYYGGTTYNRKFKGK ATLTVDKSSST AYMQLKSLISEDSAVYYCARSVGPMDYWGQGTSVTVSSDQEKSCDKTHTSPSTAPELLGGFSVFLFPFRKPDTLMVSTPEVTCVVVDVSHEDPEVKFNWYVDGVEVH NAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAK GQPREPQYYTLPPSRDELTKRQVSLTCLVKGFYPSDLAVEWESNGQPENNYKTTPP VLDSDGSFFLYSKLTVDKSRWOGGNVFSCSVMHEALHNHYTOKSLSLSFGK

# 53. G28-1 scFv VH L11S (CSC-S)H WCH2 WCH3

Nucleotide sequence:

aagettgeegeeatggtateeaeageteagtteettgggttgetgetgetgetgettaeaggtggeagatgtgaeateeagatgaete 20 agtetecageeteeetatetgeatetgtgggagagaetgteaceateaeatgtegaaeaagtgaaaatgtttaeagttatttggettggt atcagcagaaacagggaaaatctcctcagctcctggtctcttttgcaaaaaccttagcagaaggtgtgccatcaaggttcagtggca gtggatcaggcacacagttttctctgaagatcagcagcctgcagcctgaagattctggaagttatttctgtcaacatcattccgataat cgtcagcggtccagctgcagcagtctggacctgagtcggaaaagcctggcgcttcagtgaagatttcctgcaaggcttctggttact 25 cattcactggctacaatatgaactgggtgaagcagaataatggaaagagccttgagtggattggaaatattgatccttattatggtggt actacctacaaccggaagitcaagggcaaggccacaftgactgtagacaaatcctccagcacagcctacatgcagctcaagagtct gacatetgaggactetgeagtetattactgtgeaagateggteggeeetatggactactggggteaaggaaceteagteacegtete ttctgatcaggagcccaaatcttgtgacaaaactcacacatctccaccgtgctcagcacctgaactcctgggtggaccgtcagtcttc ctetteccccaaaacccaaggacaccctcatgateteccggacccetgaggtcacatgegtggtggtggtggacgtgagccacgaag 30 accetgaggteaagtteaactggtacgtggaeggegtggaggtgcataatgceaagaeaaageegeggggaggaggagtacaac agcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggagtacaagtgcaaggtctccaaca aageceteecagececeategagaaaacaatetecaaagecaaagggeageecegagaaceacaggtgtacaceetgeececa tcccgggatgagctgaccaagaaccaggtcagcctgactgcctggtcaaaggcttctatccaagcgacatcgccgtggagtgg gagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacggtccttcttctctctacagcaag 35 ctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggctctgcacaaccactacacgc agaagagceteteeetgteteegggtaaatgatetaga

### Amino acid sequence:

- MYSTAOFLGILLLWLTGGRCDIOMTOSPASLSASVOETVTITCRTSENVYSYLAW
  40 YQQKQKSPQLLVSFAKTLAEGVPSRFSGSGSGTQFSLKISSLQPEDSGSYFCQHIS
  DNPWTFGGGTELEIKGGGSGGGGSGGGGSSAVQLQQSGPESEKPGASVKISCKA
  SGYSFTGYNMNWVKQNNGKSLEWIGNIDPYYGGTTYNRKFKGKATLTVDKSSST
  AYMQLKSLTSEDSAVYYCARSVGPMDYWGGGTSVTVSSDQEFKSCDKTHTSPPC
  SAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEV
- 45 HNAKTKPREEQYNSTYRVVSVLTVLHODWLNGKEVKCKVSNKALPAPIEKTISKA KGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPENNYKTTP FVLDSDGSFFLYSKLTVDKSR WQQGNVFSCSVMHEALHNHYTQKSLSLSPGK

### 50 54. G28-1 scFv VH L11S (SSC-P)H WCH2 WCH3

Nucleotide sequence:

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WO 2005/017148 PCT/US2003/041600

aagettgeegecatggtateeaeageteagtteettgggttgetgetgetgetggettaeaggtggeagatgtgaeateeagatgaete atcagcagaaacagggaaaatctcctcagctcctggtctcttttgcaaaaaccttagcagaaggtgtgccatcaaggttcagtggca gtggatcaggcacacagttttctctgaagatcagcagcctgcagcctgaagattctggaagttatttctgtcaacatcattccgataat cg t cag cgg te cag ctg cag cag tetggac ctg ag teggaa aa ag cet t gg cget te ag t gaa gat tte ctg caa gget tetgg ta ctgg cag consideration of the consideractacctacaaccggaagttcaagggcaaggccacattgactgtagacaaatcctccagcacagcctacatgcagctcaagagtct gacatetgaggactetgeagtetattactgtgeaagateggteggeeetatggactactggggteaaggaaceteagteaeegtete ttetgateaggageecaaatettetgacaaaaetcacacateeccacegtgeecageacetgaacteetggggggacegteagtett cetetteececcaaaacccaaggacaccctcatgateteecggacccetgaggteacatgegtggtggtggacgtgagccacgaa gaccetgaggteaagtteaactggtacgtggacggegtggaggtgeataatgeeaagaeaaageegegggaggaggagtaeaa cagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggagtacaagtgcaaggtctccaac aaagccctcccagcccccatcgagaaaacaatctccaaagccaaagggcagccccgagaaccacaggtgtacaccctgcccc gagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacggctccttcttcctctacagcaag ctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaatgatctaga

# 20 Amino acid sequence:

Aminio acid sequence:
MVSTAQPLGLLLLWLTGGRCDIQMTQSPASLSASVGETVTITCRTSENVYSYLAW
YQQKQGKSPQLLVSFAKTLAEGVPSRFSGSGSGTQFSLKISSLQPEDSGSYFCQHHS
DNPWTFGGGTELEIKGGGGSGGGGSGGGGSSAVQLQQSGPESEKPGASVKISCKA
SGYSFTGYNMNWVKQNNGKSLEWIGNIDPYYGGTTYNRKFKGKATLTVDKSSST
AYMQLKSLTSEDSAVYYCARSVGPMDVWGQGTSVTVSSDQEFKSSDKTHTSPPCP
APELLGGSVFLPPPRKDELTKPFPUTCVVVDVSHEDDEVKFNWYVDGVEVH
NAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAK
GQPREPQYYTLPPSRDELTKRQVSLTCLVKGFYPSDLAVEWESNGQPENNYKTTPP
VLDSDGSFFLYSKLTVDKSRWQGGNVFSCSVMHEALHNHYTQKSLSLSFG

# II. 54. HCTLA4 HIGG1 (SSS-S)H P238SCH2 WCH3

### Nucleotide sequence:

### Amino acid sequence:

MACLGFQRHKAQLNLAARTWPCTLLFFLLFIPVFCKAMHVAQPAVVLASSRGIAS FVCEYASPGKATEVRVTVLROADSOVTEVCAATYMTGNELTFLDDSICTGTSSGN

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QVNI.TIQGLRAMDTGLYICKVELMYPPPYYLGIGNGTQIYVIDPEPCPDSDQPKSSD
KTHTSPPSSAPELLGGSSVFLFPPKPKDTLMISRTPEVTCVVDVSHEDPEVKFNWY
VDGYEVINAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPI
EKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPE
NNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSL
SPGK

# 55. Fc2-2 VL

# Nucleotide sequence:

10 gtigttaagctigecgecatggatteacaggeccaggtettatgttactgetgetatgggtactggtactgtgggacattgtgdt 
teacagtetecatectecetagetigtgagtiggaggaaggtitetatggtetggaagtecagteaggectitatataatcaeaat 
caaaagaactacttggeetggtaccageaggtaccagggeagtetectaaactgetgatttactgggcatecatagggaatetg 
ggtoctgategettecaaggeagtggatetgggacaggttgaagtetgaagagtgaaggetgaagactgaagactgaagactgaagactgagattattactgtcaagaatattatacetatectecacgtfeggaggtggaccaaagctggaaataaaaaggtggeggtggetgggggggg

15 gtgggtcgggtggcgggggggctcg

Amino acid sequence:

MDSQAQVLMILLLLWVSGTCGDIVMSQSPSSLAVSVGEKVSMSCKSSQSLLYNHN QKNYLAWYQQIPGQSPKLLIYWASTRESGVPDRFTGSGSGTDFTLTISRVKAEDLA VYYCOQYYTYPPTFGGGTKLEKGGGGSGGGSGGGGSG

# 56. FC2-2VH

20

Nucleotide sequence:

# 30 Amino acid sequence:

GSSQVQLKEŠGPGLVAPSQSLSITCTVSGFSLTVYGVNWVRQPPGKGLDWLGMIW GDGSTDYNSALKSRLSISKDNSKSQVFLKMDSLQTDDTARYYCARDHYGTHYAM DYWGGGTSVTVSSDQ

### 35 57. FC2-2scFv

Nucleotide sequence:

gtigtnagctigcegecatggalicacaggecoaggitettatgitactgetgetatggalatetggtacetgtgggacattgfgatg cacagtotecateeteetetgatgtgetgetgggaggaggaggtiletatgactggaetgeagtecagteaggecittatataatacacaat caaaagaactacttggeetggtaccageagptaceagggeagteteetaaactgetgattactgggcatecacatagggaatetgg

- 45 gttttettaaaaatggacagtetacaaactgatgacacagecaggtactactgtgccagagatcactatggtacccactatgetatgg actactggggtcaaggaacctcagtcaccgtetectetgatcag

# Amino acid sequence:

MISQAQVLMLLLLWYSGTCGDIYMSQSPSSLAVSVGEKVSMSCKSSQSLLYNIN QKNYLAWYQQIPGQSPKLLIYWASTRESGVPDRFTGSGSGTDFTLTISRVKAEDLA VYYCQQYYTYPPTFGGGTKLEIKGGGGSGGGGSGGGGSSQVQLKESGPGLVAPSQ

10

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SLSITCTVSGFSLTVYGVNWVRQPPGKGLDWLGMIWGDGSTDYNSALKSRLSISK DNSKSQVFLKMDSLQTDDTARYYCARDHYGTHYAMDYWGOGTSVTVSSDO

# 58. FC2-2 VHL11S

5 Nucleotide sequence:

gggactoteaggtgeagttgaaggagteaggactggtegggegeocteacagagectglecateacatgeacggloteg ggttetattaacegtotatggtgttaactgggttegcageotecaggaaagggttggactggetggaatgatatggggggtgt gaagcacagactataatteagctoteaaatcagactgagcatcagtaaggacaactocaagagccaagttitetaaaaatggaca gtotacaaactgatgacaagcaaggaatactgtgccagagatcactatggtaccacatatgctatggactactgggggtcaaggaa cotcagtcacogtotectgatasa

Amino acid sequence:
(GSS)QVQLKESGPGSVAPSQSLSITCTVSGFSLTVYGVNWVRQPPGKGLDWLGMI
WGDGSTDYNSALKSRLSISKDNSKSQVFLKMDSLQTDDTARYYCARDHYGTHYA
MDYWGOGTSVTVSSDO

### FC2-2 VH L11S scFv

Nucleotide sequence:

# 30 Amino acid sequence:

MDSQAQVLMILLLWVSGTCGDIVMSQSPSSLAVSVGEKVSMSCKSSQSLLYNHN QKNYLAWYQQIPGQSPKLLIYWASTRESGVPDRFTGSGSGTDFTLTISRVKAEDLA VYYCQQYYTYPPTFGGGTKLEKGGGGSGGGGSGGGSSGGGSSYQLKESGGFSVAPSQ SLSTICTVSGFSLTVYGVNWVRQPPGKGLDWLGMIWGDGSTDYNSALKSRLSISK

35 DNSKSQVFLKMDSLQTDDTARYYCARDHYGTHYAMDYWGQGTSVTVSSDQ

# 60. FC2-2 (SSS-S)H WCH2 WCH3

Nucleotide sequence:

gttgttaagettgeegecatggatteacaggeceaggttettatgttaetgetgetatgggtatetggtacetgtggggacattgtgatg 40 teacagtetecatectecetagetgtgteagttggagagaggtttetatgagetgeaagteeagteeagteagageettttatataateaeaatcaaaagaactacttggcctggtaccagcagataccagggcagtctcctaaactgctgatttactgggcatccactagggaatctgg ggtccctgatcgcttcacaggcagtggatctgggacagatttcactctcaccatcagcagagtgaaggctgaagacctggcagttta ttactgtcagcaatattatacctatcctcccacgttcggaggtggcaccaagctggaaataaaaggtggcggtggctcgggcggtg gtgggtcgggtggcgggggggctctcaggtgcagttgaaggagtcaggacctggcctggtggcgccctcacagagcctgtcc 45 gaatgatatggggtgatggaagcacagactataattcagctctcaaatccagactgagcatcagtaaggacaactccaagagccaa gttttettaaaaatggacagtetacaaactgatgacacagccaggtactactgtgccagagatcactatggtacccactatgctatgg cage acctgaact cet gggt ggaccg teag tet teet et te ee cea aaacee aa ggacacee te at gate te ee ggaccee t gag acceet gag50 gtcacatgcgtggtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataat gccaagacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggct

PCT/US2003/041600

gaatggeaaggagtacaagtgeaaggtetecaacaaageceteceagececategagaaaaceatetecaaagecaaaggge ageceegagaaccacaggtgtacaceetgeececatecegggaatgagetgacaaggaacaagtagagetgaceetgacetgeetgeetgeaaggeettetatecaaggecaategeegtggaggagaageaatggeaggeeggagaacaactacaagaccacgeetecegtg atggaeteegacggeteetteteetaagecaageteacggtggaagaagaagaagagaggggaagaagggggaagtettetaatgeteeggeetggaagaggggaaggetetgaaggetetgeaagagetggaagagggggaagatgatetagaa

Amino acid sequence:

MDSQAQVLMLLLLWVSGTCGDIVMSQSPSSLAVSVGEKVSMSCKSSQSLLYNHN
QKNYLAWYQQIPGQSPKLLIYWASTRESGVPDRFIFGSGSTDFITLTISRVKAEDLA
VYYCQQYYTYPPTFGGGTKLEIKGGGGSGGGGSGGGGSGGGSSQVQLKESGPGLVAPSQ
SLSITCTVSGFSLTVYGVMVYRQPPGKGLDWLGMIWGDGSTDYNSALKSRLSISK
DNSKSQVFLKMDSLQTDDTARYYCARDHYGTHYAMDYWGQGTSVTVSSDQEPK
SSDKTHTSPPSSAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKF
NWYVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKA
LPAPIEKTISKAKGQPRPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESN
GQPENNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQ
KSLSLSFGK

# 61, FC2-2 VHL11S (SSS-S)H WCH2 WCH3

20 Nucleotide sequence:

gttgttaagettgeegeeatggatteacaggeeeaggttettatgttaetgetgetatgggatatetggtaeetgtggggaeattgtgatgtcacag tctccatcctccctag ctgtgtcag ttggagagagaggtttctatgag ctgcag tccag tcagagccttttatataatcacaatcaa aaga acta cttggcctggtaccag caga taccag ggcag tctcctaa actgctg atttactgg gcatccactag ggaat ctggggtccctgatcgcttcacaggcagtggatctgggacagatttcactctcaccatcagcagagtgaaggctgaagacctggcagttta ttactgtcagcaatattatacctatcctcccacgttcggaggtggcaccaagctggaaataaaaggtggcggtggctcgggcggtg 25 gtgggtgggggggggggggggggtctcaggtgcagttgaaggagtcaggacctggctcggtggcgccctcacagagcctgtcc gaatgatatggggtgatggaagcacagactataattcagctctcaaatccagactgagcatcagtaaggacaactccaagagccaa gttttcttaaaaatggacagtctacaaactgatgacacagccaggtactactgtgccagagatcactatggtacccactatgctatgg actactggggtcaaggaacctcagtcaccgtctcctctgatcaggagcccaaatcttctgacaaaactcacacatccccaccgtcct 30 cagcacctgaactcctgggtggaccgtcagtcttcctcttccccccaaaacccaaggacaccctcatgatctcccggacccctgag gtcacatgcgtggtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataat gccaagacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggct gaatggcaaggagtacaagtgcaaggtctccaacaaagccctcccagcccccatcgagaaaaccatctccaaagccaaagggc 35 aaggettetateeaagegacategeegtggagtgggagageaatgggeageeggagaacaactacaagaccaegeeteeegtg ctggactccgacggctccttcttcctctacagcaagctcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctc egtgatgeatgaggetetgeacaaceactacaegeagaagageeteteeetgteteegggtaaatgatetaga

40 Amino acid sequence:

MDSQAQVLMLLLLWVSGTCGDIVMSQSPSSLAVSVGEKVSMSCKSSQSLLYNIN
QKNYLAWYQQPGQSPKLLIVWASTIRESGVPDRFTGSGSGTDFTLTISRVKAEDLA
VYYCQQYTYPPFTGGGTKLEIKGGGGSGGGSGSGGSGSQVQLKESGPGSVAPSQ
SLSITCTVSGFSLTVYGVNWVRQPPGKGLDWLGMWGDGSTDYNSALKSRLSISK
DNSKSQVFLKMDSLQTDDTARYYCARDHYGTHYAMDYWGQGTSVTVSSDQEPK
SSDKTHTSPPSSAPELLGGPSVFLPPFKPKDTLMISRTPEVTCVVDVSHEDPEVKF
NWYVDGVBVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKA
LPAPIEKTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWSN
GQPENNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHBALHNHYTG

50 KSLSLSPGK

PCT/US2003/041600

### 62. UCHL-1 VH

Nucleotide sequence:

atggg cagget tact tett catteet get act gatt gtteet ge at at gteet et ee cag attact et gaa a gagt et gge eet gggat et the state of thegcagccctcccagaccctcagtctgacttgttctttctctgggttttcactgaccacttatggtataggagtaggttggattcgtcagcct ccagggaagggtctggagtggctgacacacatttggtggaatgataataagtactataacacaggcctgaggaggccggctcacaa tetecaaggatteeteeaacaaccaagtaeteeteaagategeeaatgtggacaetgeagataeegeeacataetaetgtetetaeg gctacacttactggggccaagggactetggtcactgtctctgca

# Amino acid sequence:

10 MGRLTSSFLLLIVPAYVLSQITLKESGPGILQPSQTLSLTCSFSGFSLTTYGIGVGWIR QPPGKGLEWLTHIWWNDNKYYNTALRSRLTISKDSSNNOVLLKIANVDTADTAT YYCLYGYTYWGQGTLVTVSA

# 63. UCHL-1 VL

15 Nucleotide sequence:

> atgaagttgcctgttaggctgttggtgctgatgttctggattcctgcttccatcagtgatgttgtgatgacccaaactccactctccctgc ctgtcagtcttggagatcaggcctccatctcttgcagatctagtcagagccttctttacagtaatggaaacacctatttacattggtacct gcagaagccaggccagtctccaaaactcctgatctacaaactttccaaccgattttctggggtcccagacaggttcagtggcagtgg at cagggac agattt cacact caa gat cag cag ag t gagg ct gaggat ct gggagtt tattt ct get et caa ag tacacat gt t cggaggat ct gggaggt tattt ct get et caa ag tacacat gt t cggaggat ct gggaggt tattt ct get et caa ag tacacat gt t cggaggat ct gggaggt tattt ct get et caa ag tacacat gt t cggaggat ct gggaggt tattt ct get et caa ag tacacat gt t cggaggat ct gggaggt tattt ct get et caa ag tacacat gt t cggaggat ct gggaggt tattt ct get et caa ag tacacat gt t cggaggat ct gggaggt tattt ct get et caa ag tacacat gt t cggaggat ct gggaggat ggacgttcggtggaggcaccaagctggaaatcaaa

20

# Amino acid sequence:

MKLPVRLLVLMFWIPASISDVVMTQTPLSLPVSLGDQASISCRSSQSLLYSNGNTYL HWYLQKPGQSPKLLIYKLSNRFSGVPDRFSGSGSGTDFTLKISRVEAEDLGVYFCS 25 OSTHVPWTFGGGTKLEIK

#### 64. UCHL-1 scFv

Nucleotide sequence:

gitgttaagettgeegecatgaagitgeetgttaggetgttggtgatgttetggatteetgetteeateagtgatgttgtgatgaeee 30 aaactecaeteteectgeetgteagtettggagateaggeeteetetttgeagatetagteagageettetttaeagtaatggaaae teaaagtacacatgtteegtggacgtteggtggaggcaccaagetggaaateaaagatggeggtggteggteggtggtggatet ggaggaggtgggagcteteagattactetgaaagagtctggeeetgggatettgeageeeteeeagaeeetcagtctgacttgttett 35 tctctgggttttcactgaccacttatggtataggagtaggttggattcgtcagcctccagggaagggtctgagagtggctgacacacat tiggtggaatgataataagtactataacaagccctgaggagccggctcacaatctccaaggattcctccaacaacaacaagtactcct caagatcgccaatgtggacactgcagataccgccacatactactgtctctacggctacacttactggggccaagggactctggtca ctgtctctgctgatca

#### 40 Amino acid sequence:

MKLPVRLLVLMFWIPASISDVVMTOTPLSLPVSLGDOASISCRSSOSLLYSNGNTYI. HWYLQKPGQSPKLLIYKLSNRFSGVPDRFSGSGSGTDFTLKISRVEAEDLGVYFCS QSTHVPWTFGGGTKLEIKDGGGSGGGGGGGGGGSOITLKESGPGILOPSOTLSLTCS FSGFSLTTYGIGVGWIRQPPGKGLEWLTHIWWNDNKYYNTALRSRLTISKDSSNN

45 QVLLKIANVDTADTATYYCLYGYTYWGQGTLVTVSAD

### 65. UCHL-1 VH I11SL12S

Nucleotide sequence:

50 gggageteteagattaetetgaaagagtetggeeetgggatettgeageeeteeagaeeeteagtetgaettgttettetetgggttt tcactgaccacttatggtataggagtaggttggattcgtcagectccagggaagggtctggagtggctgacacacatttggtggaat

gataataagtactataacacagccotgaggagcoggotcacaatotcoaaggattoctocaacaaccaagtactoctoaagatogc caatytgacactgoagatacogocacatactactgtotctacggotacacttactggggccaagggactotggtcactgtototget satea

5 Amino acid sequence:

(GSS)QITLKESGPGSSQPSQTLSI.TCSFSGFSLTTYGIGVGWIRQPPGKGLEWLTHW WNDNKYYNTALRSRLTISKDSSNNQVILKIANVDTADTATYYCLYGYTYWGQGT LVTVSAD

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15

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# 66. UCHL-1 scFv VH L11S

Nucleotide sequence:

Amino acid sequence:

25 MKLPVRLLVLMFWIPASISDVVMTQTPLSLPVSLGDQASISCRSSQSLLYSNGNTYL
HWYLQRPGQSPKLLIYKLSNRFSGVPDRFSGSGSGTDFTLKISRVBAEDLGVYFCS
QSTHVPWTFGGGTKLEIKDGGGSGGGSGGGGSGGTLKESGPGSSQPSQTLSLTC
SFSGFSLTTYGIGVGWIRQPPGKGLEWLTHIWWNDNKYYNTALRSRLTISKDSSNN
OVLKIANVDTADTATTYYCLYGYTYWGGGTLVTVSAD

30

# 67. UCHL-1 scFv (SSS-S)H WCH2 WCH3

Nucleotide sequence:

gttgttaagettgeegeeatgaagttgeetgttaggetgttggtgetgatgttetggatteetgetteeateagtgatgttgtgatgaeee35 aaactccactctccctgcctgtcagtcttggagatcaggcctccatctcttgcagatctagtcagaggccttctttacagtaatggaaac caggtt cagtgg cagtgg at cagggacag at ttc acact caa gat cag cag agt gg agg ctg agg at ctg gg agt tt at ttc tgc tc agg transfer of the contract can be caused as the contract can be contracted as the contcaaagtacacatgttccgtggacgttcggtggaggcaccaagctggaaatcaaagatggcggtggctcgggcggtggtggatct ggaggaggtgggagctctcagattactctgaaagagtctggccctgggatcttgcagccctcccagaccctcagtctgacttgttctttototgggttttcactgaccacttatggtataggagtaggttggattcgtcagcctccagggaagggtclggagtggctgacacacat40 ttggtggaatgataataagtactataacacagccctgaggagccggctcacaatctccaaggattcctccaacaaccaagtactcct caagategeeaatgtggacaetgeagataeegeeacataetaetgtetetaeggetaeaettaetggggeeaagggactetggtea ctgtctctgetgatcaggagcccaaatcttctgacaaaactcacacatccccaccgtcctcagcacctgaactcctgggtggaccgtcagtetteetetteeeccaaaaacceaaggacacceteatgateteeeggacceetgaggteacatgegtggtggtggaegtgage 45 cacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagccgcggggaggagca gtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggagtacaagtgcaaggtc tecaacaaageceteecagececcategagaaaaccatetecaaagecaaagggcagececgagaaccacaggtgtacaccet gagtgggagagcaatgggcagccggagaacaactacaagaccacgceteccgtgctggactccgacggeteettetteetetae 50  $agca agcte accept {\tt ggacaagagcaggt} {\tt ggcagcaggggaacgtett} {\tt ctcatgctccgt} {\tt gatgcatgaggctctgcacaaccact}$ acaegeagaagageeteteeetgteteegggtaaatgatetaga

WO 2005/017148 PCT/US2003/041600

Amino acid sequence:

MKLPVRLLVÍLMEWIPASISDVVMTOTPLSILPVSLGDQASISCRSSQSLLYSNGNTYL
HWYLQKPGQSPKLLIYKLSNRFSGVPDRFSGSGSGTDFTLKISRVEAEDLGYYFCS
QSTHVPWTFGGGTKLEIKDGGSSGGGSGSGGGSSQITLKESGPGILQPSQTISLTCS
FSGFSLTTYGIGVGWIRQPPGKGLEWLTHIWWNDNKYYNTALRSRLTISKDSSNN
QVLLKIANVDTADTATYYCLYGYTYWGQGTLVTVSADQEPKSSDKTHTSPPSSSN
ELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVFVHNA
KTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIFKTISKAKGQ
PREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWSNGQPENNYKTTPPVL
DSDGSFFLYSKLTVDKSRWOQGNVFSCSVMHBALHNHYTOKSLSLSPGK

### 68. UCHL-1 scFv VHL11S (SSS-S)H WCH2 WCH3

15 Nucleotide sequence:

gttgttaagettgeegeeatgaagttgeetgttaggetgttggtgetgatgttetggatteetgetteeateagtgatgttgtgatgacee aaactccactetccctgcctgtcagtcttggagatcaggcctccatctcttgcagatctagtcagagccttctttacagtaatggaaac acctatttacattggtacctgcagaagccaggccagtctccaaaactcctgatctacaaactttccaaccgattttctggggtcccaga cagettcagtggcagtggatcagggacagatttcacactcaagatcagcagagtggaggctgaggatctgggagtttatttctgctc 20 ggaggaggtgggagctctcagattactctgaaagagtctggccctgggagctcccagccctcccagaccctcagtctgacttgttc tttetetgggtttteaetgaceaettatggtataggagtaggttggattegteageeteeagggaagggtetggagtggetgaeaeae attiggtggaatgataataagtactataacacagccctgaggagccggctcacaatctccaaggattcctccaacaaccaagtactc cteaagategeeaatgtggacactgeagatacegeeacatactactgtetetaeggetaeacttaetggggeeaagggactetggte 25 actgtetetgetgateaggageceaaatettetgacaaaacteacacatececacegteeicagcacetgaacteetgggtggaceg teagtetteetetteeceeaaaacccaaggacaccteatgateteeeggacccetgaggteacatgegtggtggacgtgag ccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcatautgccaagacaaagccgcgggaggagga agtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggagtacaagtgcaaggt ctccaacaaagccctcccagcccccatcgagaaaaccatctccaaagccaaagggcagccccgagaaccacaggtgtacaccc 30 ggagtgggagagcaatgggcagccggagaacaactacaagaccacgcctcccgtgctggactccgacggctccttcttcctcta cagcaagetcaccgtggacaagagcaggtggcagcaggggaacgtcttctcatgctccgtgatgcatgaggctctgcacaacca ctacacgcagaagagcctctccctgtctccgggtaaatgatctagaa

35 Amino acid sequence:

MKLPVRLLVÍMFWPASISDVVMTOTPI SLPVSLGDQASISCRSSQSILYSNGNTYL
HWYLQKPGQSPKLLIYKLSNRFSGVPDRPSGSSGSGTDFTLKISRVEAEDLGVYFCS
QSTHVPWTFGGGTKLEIKDGGGSGGGGSSGGGGSSQITLKESGPGSSQPSQTTLSLTC
SFSGFSLTTYGIGVGWIRQPFCKGLEWLTHIWWNDNKYYNTALRSRLTISKDSSNN
40 QVLLKIANVDTADTATYYCLYGYTYWGGGTLVTVSADQEPKSSDKTHTSPPSSAP
ELLGGPSVFLFPFKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYVDGVEVHNA
KTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIEKTISKAKGQ
PREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWSNGQPENNYKTTPPVL

69, 5B9 VH L11S

45

Nucleotide sequence:

DSDGSFFLYSKLTVDKSRWOOGNVFSCSVMHEALHNHYTOKSLSLSPGK

tgcaacctaatgacacagccatttattactgtgccagaaatgggggtgataactacccttattactatgctatggactactggggtcaaggaacctcagtcaccgtctcctcag

Amino acid sequence:

5 (GSS)QVQLK\(\)GSPGSV\(\)QSS\(\)QSS\(\)QSS\(\)QSS\(\)QSS\(\)QSS\(\)QSS\(\)QSS\(\)QSS\(\)QSS\(\)SSS\(\)SSS\(\)CTYAVHWVR\(\)QSPGKGLEWLGVI WSGGTIDYNAAFISRLSITKDDSKS\(\)QVFKMNSL\(\)QPNDTAIYYCARNGGDNYPYY YAMDYWGOGTSVTVSS

# 10 73. 5B9 VH L11S scFv

Nucleotide sequence:

Amino acid sequence:

MRFSAQLLGLLVLWIPGSTADIVMTQAAFSNPVTLGTSASISCRSSKSLLHSNGITY
LYWYLQKPGQSPQLLIYQMSNLASGVPDRFSSSGSGTDFTLRISRVEAEDVGVYYC
AQNLELPLIFGAGTKLELKRGGGGSGGGGSSQVQLKSGVGPGSVQSSQSLSI
TCTVSGFSLTTYAVHWVRQSPGKGLEWLGVIWSGGITDYNAAFISRLSITKDDSKS
QVFFKMNSLQPNDTAIYYCARNGGDNYPYYYAMDYWGQGTSVTVSS

30

15

20

# 70. 5B9 scFv VHL11S (SSS-S)H WCH2 WCH3

Nucleotide sequence:

 $a a gett {\tt geogecat} gaggt {\tt tetg} gett {\tt etg} gg {\tt gett} get {\tt tetg} gate {\tt cetg} gate {\tt cetg} gate {\tt cactg} cagat {\tt attg} t {\tt gate} gac {\tt gett} gate {\tt cetg} gate {\tt cetg} gate {\tt cactg} cagat {\tt attg} t {\tt gate} gac {\tt gate} gate {\tt cetg}  ggetgeatteteeaateeagteactettggaacateagetteeateteetgeaggtetagtaagagteteetaeatagtaatggeatea 35 cttatttgtattggtatctgcagaagccaggccagtctcctcagctcctgatttatcagatgtccaaccttgcctcaggagtcccagaca  ${\tt ggttcagtagcagtgggtcaggaactgatttcacactgagaatcagcagagtggaggctgaggatgtgggtgtttattactgtgctc}$ cgggtggcggcgggagctctcaggtgcagctgaagcagtcaggacctggctcagtgcagtcctcacagagcctgtccatcacct gcacagtctctggtttctcattaactacctatgctgtacactgggttcgccagtctccaggaaagggtctggagtggctgggagtgat40 atggagtggtggaatcacagactataatgcagctttcatatccagactgagcatcaccaaggacgattccaagagccaagttttctttaaaatgaacagtctgcaacctaatgacacagccatttattactgtgccagaaatgggggtgataactacccttattactatgctatgga ctactggggtcaaggaacctcagtcaccgtetcctctgatcaggagcccaaatcttctgacaaaactcaccaccatccccaccgtcctcageaectgaactcetgggtggaccgtcagtetteetetteececcaaaacecaaggacacectcatgateteecggaccectgagg tcacatgcgtggtggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgcactggtgaggtgcataatgcactggacgtggacgtggaggtgcataatgcactggacgtggacgtggaggtgcataatgcactggacgtggacgtggacgtggacgtgaggtgcataatgcactggacgtgaggtgaaggaagaaggaagaaggaaggaagaaggaagaaggaaggaagaaggaaggaagg45 caagacaaagccgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctga alggcaaggagtacaagtgcaaggtetecaacaaageceteecagececeategagaaaaccatetecaaagecaaagggcag ccccgagaaccacaggtgtacaccctgccccatcccgggatgagetgaccaagaaccaggtcagcctgacctgcctggtcaaa ggettetatecaagegacategeegtggagtgggagageaatgggeageeggagaacaactacaagaecaegeeteeegtget ggactccgacggctccttcttctctctacagcaagctcaccgtggacaagagcaggtggcagcaggagggaacgtcttctcatgctcc 50 gtgatgcatgaggctctgcacaaccactacacgcagaagagcctctccctgtctccgggtaaatgatctagag

Nucleotide sequence: aagettgccgccatggattttcaagtgcagattttcagcttcctgctaatcagtgcttcagtcataattgccagaggacaaattgttctct50 cccag tete cag caate ct g to tge at ctc cag g g g a g a a g g tea caat g a ctt g cag g g c cag ctca a g t g taa g t ta cat g caet g cag g c cag ctca a g t g taa g t ta cat g caet g c

#### 2H7 scFv VH L11S (SSS-S)H WCH2 WCH3 78.

FNPPTFGAGTKLELKDGGGSGGGGSGGGSSQAYLQQSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKQTPRQGLEWIGAIYPGNGDTSYNQKFKGKATLTVDKSSS 40 TAYMQLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDOEPKSSDK THTSPPSSAPELLGGSSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV DGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPAPIE KTISKAKGQPREPQVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQPEN NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS 45 PGK

QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCQOWS

Amino acid sequence: MDFQVQIFSFLLISASVIIARGQIVLSQSPAILSASPGEKVTMTCRASSSVSYMHWY

Amino acid sequence:

tacacatttaccagttacaatatgcactgggtaaagcagacacctagacagggcctggaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggattggagctatttatccaggaaatggattggagctatttatccaggaaatggattggattggagctatttatccaggaaatggattggattggattggagctatttatccaggaaatggattggattggagctatttatccaggaaatggattggggtgatacttectacaatcagaagttcaagggcaaggccacactgactgtagacaaatcctccagcacagcctacatgcagctcag 25 . agggaccaeggteaccgtetettetgateaggageceaaatettetgacaaaaeteaeacateeecaccgteeteageacetgaaet cetgggggategteagtetteetetteececcaaaacccaaggacacceteatgateteeeggacccetgaggteacatgcgtgg . 'tggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagc egegggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggag tacaagtgcaaggtctccaacaaagccctcccagcccccatcgagaaaacaatctccaaagccaaagggcagccccgagaacc 30 agegacategeegtggagtgggagageaatgggeageeggagaacaactacaagaceaegeeteeegtgetggacteegaeg geteettetteetetacageaageteacegtggacaagageaggtggcageaggggaaegtetteteatgeteegtgatgeatgag getetgeacaaceaetacaegeagaagageeteteeetgteteegggtaaatgatetaga

ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg gcagtgggtctgggacctcttactctctcacaatcagcagagtggaggctgaagatgctgccacttattactgccagcagtggagttt 20 ggagetete aggettateta cage agtet gggget gagte ggt gagge et ggggeet cagt gaag at gteet geaagget tet ggegetet ggggeet gagget gaggetet ggggeet gagget 
Nucleotide sequence: a a gett geogecat gg att tt caagt ge a gat tt te agette et getaat eagt gette agt eat aatt gecag ag gae aa att gete et getaat en gete gedeel gedoel gedoel gedeel gedoel gedoel gedoel gedoel gedoel gedoel gedoelcccagtctccagcaatcctgtctgcatctccaggggagaaggtcacaatgacttgcagggccagctcaagtgtaagttacatgcact

ENNYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLS 10 LSPGK

2H7 scFv VH L11S (SSS-S)H P238SCH2 WCH3

MRFSAQLLGLLVLWIPGSTADIVMTQAAFSNPVTLGTSASISCRSSKSLLHSNGITY LYWYLQKPGQSPQLLIYQMSNLASGVPDRFSSSGSGTDFTLRISRVEAEDVGVYYC AONLELPLTFĞAGTKLELKRGGGGSGGGGGGGGGSSOVQLKQSGPGSVOSSQSLSI TCTVSGFSLTTYAVHWVRQSPGKGLEWLGVIWSGGITDYNAAFISRLSITKDDSKS OVFFKMNSLOPNDTAIYYCARNGGDNYPYYYAMDYWGQGTSVTVSSDQEPKSS DKTHTSPPSSAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNW YVDGVEVHNAKTKPREEQYNSTYRVVSVLTVLHQDWLNGKEYKCKVSNKALPA PIEKTISKAKGOPREPOVYTLPPSRDELTKNQVSLTCLVKGFYPSDIAVEWESNGQP

WO 2005/017148 PCT/US2003/041600

15 76.

ggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg geagtgggtetgggaeetettacteteteaeaateageagagtggaggetgaagatgetgeeacttattactgeeageagtggagttt taacccaccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggtggatctggaggaggtg ggagctctcaggcttatctacagcagtctggggctgagtcggtgaggcctggggcctcagtgaagatgtcctgcaaggcttctggc tacacatttaccagttacaatatgcactgggtaaagcagacactagacagggcetggaatggattggagctatttatccaggaaat cagcctgacatctgaagactctgcggictatttctgtgcaagagtggtgtactatagtaactcttactggtacttcgatgtctggggcac agggaccaeggteaccgletettetgateaggageccaaatettetgacaaaacteacacatececaeegteeteageacetgaact cctggggggaccglcagicttcctcttcccccaaaacccaaggacaccctcatgatctcccggacccctgaggtcacatgcgtgg tggtggacgtgagccacgaagaccetgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagc cgcgggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggag tacaagtgeaaggtetecaacaaagceeteceagceeceategagaaaacaatetecaaageeaaagggeagceeegagaace acaggtgtacaccctgccccatcccgggatgagctgaccaagaaccaggtcagcctgacctgctcgtcaaaggcttctatccc agcgacategeegtggagtgggagagcaatgggeageeggagaacaactacaagaeeaegeeteeegtgetggaeteegaeg geteettetteetetacageaageteacegtggacaagageaggtggacaggggaacgletteteatgeteegtgatgeatgag getetgeacaaceactacaegeagaagageeteteeetgteteegggtaaatgatetaga

Amino acid sequence:

MDFOVOIFSFLLISASVIIARGOIVLSOSPAILSASPGEKVTMTCRASSSVSYMHWY 20 QQKPGSSPKPWIYAPSNLASGVPARFSGSGSGTSYSLTISRVEAEDAATYYCOOWS FNPPTFGAGTKLELKDGGGSGGGGSGGGSSOAYLOOSGAESVRPGASVKMSCK ASGYTFTSYNMHWVKOTPROGLEWIGAIYPGNGDTSYNOKFKGKATLTVDKSSS TAYMOLSSLTSEDSAVYFCARVVYYSNSYWYFDVWGTGTTVTVSSDQEPKSSDK THTSPPSSAPELLGGPSVFLFPPKPKDTLMISRTPEVTCVVVDVSHEDPEVKFNWYV DGVEVHNAKTKPREEOYNSTYRVVSVLTVLHODWLNGKEYKCKVSNKALPAPIE 25 KTISKAKGOPREPOVYTLPPSRDELTKNOVSLTCLVKGFYPSDIAVEWESNGOPEN NYKTTPPVLDSDGSFFLYSKLTVDKSRWQQGNVFSCSVMHEALHNHYTQKSLSLS PGK

30

35

45

10

15

# 2H7 scFv VH L11S (CSS-S)H WCH2 WCH3

Nucleotide sequence:

a agett geogecat gg att tica aget georga att tica get te ct get a at cag t get te agt cat a att georga ga a a att get te ct get a at cag t get te aget a att georga ga a a att get te ct get a att georga ga ccagic tecagca at cet glot geat et ceaggggaga aggteaca at gacttgeagggee agetea agt tacat geat the compact of the compact and the compact and the compact and the compact aggree agetea aggree agetea agt to the compact aggree agete aggree aggtaccagcagaagccaggatcctcccccaaaccctggatttatgccccatccaacctggcttctggagtccctgctcgcttcagtg geagtgggtetgggacetettaeteteteaeaateageagagtggaggetgaagatgetgeeaettattaetgeeageagtggagitt taacccacccacgttcggtgctgggaccaagctggagctgaaagatggcggtggctcgggcggtggttggatctggaggaggtg ggagcteteaggettatetacagcagtetggggetgagteggtgaggeetggggceteagtgaagatgteetgeaaggettetgge taca catttac cagttaca at at geact gggtaa ag cagacacctag ac ag ggcct ggaat ggatt ggag ct at ttat ccag gaa at the cagacactag according to the control of the cagacactag according to the cagacactag ac40 agggaccaeggteaecgtetettetgateaggageceaaatettgtgacaaaacteaeacateeeeacegteeteageacetgaaet cctggggggaccgtcagtcttcctcttcccccaaaacccaaggacacctcatgatclcccggacccctgaggtcacatgcgtgg tggtggacgtgagccacgaagaccctgaggtcaagttcaactggtacgtggacggcgtggaggtgcataatgccaagacaaagc egegggaggagcagtacaacagcacgtaccgtgtggtcagcgtcctcaccgtcctgcaccaggactggctgaatggcaaggag tacaagtgcaaggtctccaacaaagccctcccagccccatcgagaaaacaatctccaaagccaaagggcagccccgagaacc  $a caggt {\tt gtacaccet} g ceccateccegg {\tt gatgaget} g accaagaaccaggt {\tt cagect} g acct {\tt gcct} g t {\tt caaagget} t {\tt ctatccc}$ agegacategeegtggagtgggagageaatgggeageeggagaacaactacaagaccaegeetceegtgetggacteegaeg geteettetteetetacageaageteacegtggacaagageaggtggeageaggggaacgtetteteatgeteegtgatgeatgag 50 gctetgeaeaaceaetaeaegeagaagageeteteeetgteteegggtaaatgatetaga